

Phytotherapy for *Escherichia coli*: Iranian Native Medicinal Plants with Anti-*Escherichia coli* Effect

Reza Khadivi Boroujeni¹, Negar Khodabandehloo², Mahdiye Ghadirali Khorzoughi², Reza Mohammadrezaei-Khorramabadi³, Mansour Amraei⁴, Somayeh Delfani^{5*}

¹Department of Hygiene, Science and Research Branch, Islamic Azad University, Tehran, Iran

²Graduated Student of Veterinary Medicine, Faculty of Veterinary Medicine, Science and Research Branch, Islamic Azad University, Tehran, Iran

³Student Research Committees, Lorestan University of Medical Sciences, Khorramabad, Iran

⁴Biotechnology and Medicinal Plants Research Center, Ilam University of Medical Sciences, Ilam, Iran

⁵Razi Herbal Medicines Research Center, Lorestan University of Medical Sciences, Khorramabad, Iran

Available Online: 1st May, 2017

ABSTRACT

Escherichia coli bacteria as a gram-negative bacilli of Enterobacteriaceae family causes different diseases in human such as wound infection, pneumonia, urinary tract infections, meningitis, premature and weak children birth, peritonitis and cholecystitis. One of the major pathogens that has shown resistance to most antibiotics is *Escherichia coli*. Therefore, development of natural antibacterial agents such as medicinal plants for the treatment of infectious diseases is necessary. This study was aimed to present the phototherapy of Iranian native medicinal plants with anti-*Escherichia coli* effect. The required information was obtained by searching key words such as *Escherichia coli*, Native medicinal plant of Iran, medicinal plant extracts or essential oils of related published articles in authentic scientific databases. Results showed that different native medicinal plants were effective against *E. coli* in Iran, including *Cuminum cyminum*, *Mentha piperit*, *Myrtus communis*, *Mentha pulegium*, *Rosmarinus officinalis*, *Stachys inflate*, *Thymus caramanicus*, *Zataria multiflora*, *Ziziphora clinopodioides*, *Allium cepa*, *Ocium gratissimum*, *Olea europaea* and *Carum copticum*.

Keywords: Infectious diseases, *Escherichia coli*, Medicinal plants, Iran.

INTRODUCTION

Nowadays, *Escherichia coli* bacteria are major causes of infectious diseases in developing countries with high health costs and mortality. Current antibiotics are expensive and pathogenic strains resistance to them is one of the most important issues in clinical treatments¹. *E. coli* bacteria as a gram-negative bacilli of Enterobacteriaceae family causes different diseases in human such as wound infection, pneumonia, urinary tract infections (UTI), meningitis, premature and weak children birth, peritonitis and cholecystitis^{2,3}. *Escherichia coli* has been introduced as an indicator of foods fecal contamination in many international standards⁴. Recently, a number of antibiotics have lost their therapeutic effects due to development of strains resistant to antibiotics. In addition, antibiotics have various side effects such as allergic reactions, severe allergies and immunity suppression⁵. *E. coli* is one of the major pathogens that have shown resistance to most antibiotics⁶ and uses various mechanisms for resistance to beta-lactams, which include changes in outer membrane proteins, over-production of chromosomal and plasmid cephalosporinase or production of a broad spectrum beta lactamase^{7,8}. Therefore, development of natural antibacterial agents such as medicinal plants for the

treatment of infectious diseases is necessary. Medicinal plants have wide range of health effects⁸⁻¹². Furthermore, these plants other than treatment, are useful in prevention of a wide variety of diseases²⁰⁻²³. Medicinal plants have been used for a long time over generations and it is believed that they are safer than synthetic drugs²⁴⁻²⁶. Many native medicinal plants in Iran have antimicrobial and antibiotic properties. So the aim of this study is reporting medicinal plants with anti-*Escherichia coli* effect.

METHODS

The required information was obtained by searching key words such as *Escherichia coli*, native medicinal plant of Iran, medicinal plant extracts or essential oils of published articles in authentic scientific databases such as Sciedirect, Blackwell Wiley, Springer, Google scholar and Scientific Information Database (SID) and Magiran. All related articles were selected.

RESULTS

According to literature reviews, results showed that different native medicinal plants in Iran were effective against *E. coli*, including *Cuminum cyminum*, *Mentha piperit*, *Myrtus communis*, *Mentha pulegium*, *Rosmarinus*

*Author for Correspondence: somayehdelfani@gmail.com

Table 1: List of Iranian native medicinal plants effective against *E.coli*.

No	Scientific name	Family name	Description
1	<i>Cuminum cyminum</i>	Apiaceae	Results of a study showed that essential oil of <i>Cuminum cyminum</i> had the highest antimicrobial effect on <i>E. coli</i> ATCC 25922 in 1, ½, ¼ and 1/8 dilutions and growth inhibition zone diameter of this essential oil was 13 mm ²⁷
2	<i>Mentha piperita</i>	Lamiaceae	Results of a study showed antimicrobial effects of <i>Mentha piperita</i> essential oil at 1 µg/gr concentration was 3+, for 2 µg/gr was 2+ and growth inhibition zone diameter of this essential oil was 31.33 mm. D-values for this essential oil was 2.14 ²⁸
3	<i>Myrtus communis</i>	Myrtaceae	Results of a study showed that antimicrobial effects of <i>Myrtus communis</i> essential oil at 1 µg/gr concentration was 3+, for 2 µg/gr was 2+ and growth inhibition zone diameter of this essential oil was 13 mm. D-values for this essential oil was 2.8 ²⁸
4	<i>Mentha pulegium</i>	Lamiaceae	Results of an experimental study showed that MIC and MBC of <i>Mentha pulegium</i> essential oil was 4 µg/gr ²⁹
5	<i>Rosmarinus officinalis</i>	Lamiaceae	Results of the antimicrobial activity of <i>Rosmarinus officinalis</i> essential oils in a showed growth inhibition zone diameter were 16, 12.5, 9, 8 and 7 mm in 1, 1/2, 1/4, 1/8 and 1/16 dilutions, respectively ³⁰
6	<i>Stachys inflata</i>	Labiatae	Results of an experimental study showed that MIC of <i>Stachys inflata</i> essential oil was 5 µg/gr ²¹
7	<i>Thymus caramanicus</i>	Lamiaceae	Results of an experimental study showed that MIC of <i>Thymus caramanicus</i> essential oil of Vegetative stage, Floral budding, Flowering stage and seed set against <i>E.coli</i> was 0.9 µL ²²
8	<i>Zataria multiflora</i>	Lamiaceae	Results of an experimental study showed that minimum growth inhibition zone diameter of <i>Zataria multiflora</i> essential oil collected from Hajiabad, Farashabad, Yazd, Najafabad and poldokhtar area against <i>E.coli</i> were 4, 16, 8, 2 and 16 mm ³³
9	<i>Ziziphora clinopodioides</i>	Lamiaceae	Results of an experimental study showed that IZ index and MIC of <i>Ziziphora clinopodioides</i> essential oil against <i>E.coli</i> were 20±0.5 and 3.75±0.1, respectively. These indices for 1,8-Cineole were 20±0.1 and 11.68±0.5, and for Pulegone were 12±0.4 and 7.2±0.3, respectively against <i>E. coli</i> ³⁴
10	<i>Allium cepa</i>	Liliaceae	Results of an experimental study based on disc diffusion method showed that growth inhibition zone diameter of 6.25, 12.5, 25 and 50 percent concentrations of yellow onion extract were 0.9 ± 0.3, 1.8 ± 0.2, 2.3 ± 0.2 and 6.8 ± 0.5 mm, respectively and MIC and MBC of this extract were less than 50 % and 25 %, respectively ³⁵
11	<i>Allium spp.</i>	Liliaceae	Results of an experimental study based on disc diffusion method showed that growth inhibition zone diameter of 6.25, 12.5, 25 and 50 percent concentrations of Red onion extract were 0.8 ± 0.2, 2.1 ± 0.3, 3.5 ± 0.3 and 8.1 ± 0.3 mm, respectively and MIC and MBC of this extract were less than 50 % and 25 %, respectively ³⁶
12	<i>Allium spp.</i>	Liliaceae	Results of an experimental study based on disc diffusion method showed that growth inhibition zone diameter of 6.25, 12.5, 25 and 50 percent concentrations of Red onion extract were 0.4 ± 0.2, 1.4 ± 0.2, 2.2 ± 0.2 and 3 ± 0.2 mm, respectively and both MIC and MBC of this extract were less than 50 %, respectively ³⁶
13	<i>Ocimum gratissimum</i>	Lamiaceae	Results of a study showed that growth inhibition zone diameters of <i>Ocimum gratissimum</i> extract against <i>E. coli</i> were 5 mm and 9 mm in 100 mg/ml and 200 mg/ml concentrations, respectively ³⁶
14	<i>Olea europaea</i>	Oleaceae	Results of a study showed that growth inhibition rate of <i>Dezful</i> and <i>Yellow olive</i> varieties against <i>E. coli</i> were 93% and 94.9% in 20 mg/ml concentration, respectively ³⁷
15	<i>Carum copticum</i>	Asteraceae	Results of an experimental showed that growth inhibition zone diameter of <i>Carum copticum</i> extract against <i>E. coli</i> were 23 and

28 mm³⁸16 *Scrophularia Striata*

Results of an in vitro study showed positive effect of *Scrophularia Striata* extract against *E. coli*³⁹

officinalis, *Stachys inflate*, *Thymus caramanicus*, *Zataria multiflora*, *Ziziphora clinopodioides*, *Allium cepa*, *Ocium gratissimum*, *Olea europaea* and *Carum copticum* which are listed in Table 1.

DISCUSSION

Results showed that different native medicinal plants in Iran including *Cuminum cyminum*, *Mentha piperit*, *Myrtus communis*, *Mentha pulegium*, *Rosmarinus officinalis*, *Stachys inflate*, *Thymus caramanicus*, *Zataria multiflora*, *Ziziphora clinopodioides*, *Allium cepa*, *Ocium gratissimum*, *Olea europaea* and *Carum copticum* were effective against *E. coli*.

It is not clear that with what mechanisms these plants have anti *E. coli* activities. Medicinal plants have various compounds by which they may act against bacteria. Phenolic compounds are a group of compounds which are abundant in most of these and other medicinal plants⁴⁰⁻⁵⁰. These plants mostly have shown to possess antibacterial effects⁵¹⁻⁵⁸. Therefore, these compounds might be responsible for anti *E. coli* activity of these plants. More importantly these compounds in medicinal plants have antioxidant activities⁵⁸⁻⁶⁷.

Infections are associated with increase in free radicals and oxidative stress which may exacerbate the condition. Hence, medicinal plants may also be beneficial in infected diseases by their antioxidant activities and reducing oxidative stress. Oxidative stress is involved in various diseases such as neurological disorders, inflammation, ischemia/reperfusion ischemia/reperfusion, diabetes, atherosclerosis, cardiovascular diseases and wound complication⁶⁸⁻⁷⁸.

The oxidative stress involves many changes, including alterations in redox state. Antioxidants, particularly the herbal medicines with antioxidant activities are able to counteract these complications^{79,80}. Various studies have demonstrated promising results for counteracting these conditions by medicinal plant especially for life threatening diseases⁸¹⁻⁸⁴. These agents are also effective in inhibition of toxic agents induced complications. Therefore, these plants other than infections may also be beneficial in other disease.

REFERENCES

- Nakhjavani, FA, Emaneini M, Hosseini H, Taherikalani M, Mirsalehian A. Molecular analysis of typical and atypical enteropathogenic *Escherichia coli* (EPEC) isolated from children with diarrhoea. Journal of Medical Microbiology 2013; 62 (PART 2); 191-195.
- Malekzadeh F. Microbiology.2nd ed. In Tehran University press; 1994; 21- 50.
- Naderinasab M, Rashed T, Nazem M. Laboratory bacteriology In Mashhad: Imam Reza press: 1997; 24-29.
- Jay JM, Loessner MJ, Golden DA. Modern Food Microbiology, 7th edition. Springer, New York, 2016.
- Berahou A, Auhmani A, Fdil N, Benharref A, Jana M, Gadhi CA. Antibacterial activity of *quercus ilex* bark's extracts. J Ethnopharmacol 2007; 112(3): 426-429.
- Akinfogunla OJ, Eghafona NO, Ekoi OH. Diarrheal *Escherichia coli* (DEC): prevalence among in and ambulatory patients and susceptibility to antimicrobial chemotherapeutic agents. J Bact Resea 2009; 1(3): 34-38.
- Poirel L, Naas T, Guibert M, Labia R, Nordmann P. Molecular and biochemical characterization of VEB-1, a novel class A extended-spectrum β-lactamase encoded by an *Escherichia coli* integron gene. Antimicrobial agents and chemotherapy. 1999; 43(3):573-81.
- Bahmani M, Banihabib E, Rafieian-Kopaei M, Gholami-Ahangaran M. Comparison of Disinfection Activities of Nicotine with Copper Sulphate in water Containing *Limnatis nilotica*. Kafkas Univ Vet Fak Derg 2015; 21 (1): 9-11.
- Ebrahimie M, Bahmani M, Shirzad H, Rafieian-Kopaei M, Saki K. A Review Study on the Effect of Iranian Herbal Medicines on Opioid Withdrawal Syndrome. J Evid Based Complementary Altern Med. 2015; 20(4):302-9.
- Bahmani M, Shirzad H, Rafieian S, Rafieian-Kopaei M. Silybum marianum: Beyond Hepatoprotection. J Evid Based Complementary Altern Med. 2015; 20(4): 292-301.
- Ganji-Arjenaki M, Rafieian-Kopaei M. Probiotics are a good choice in remission of inflammatory bowel diseases: A Meta Analysis and systematic review. Journal of Cellular Physiology. 2017 Mar 15. PubMed PMID: 28294322.
- Baharvand-Ahmadi B, Bahmani M, Tajeddini P, Naghdi N, Rafieian-Kopaei M. An ethno-medicinal study of medicinal plants used for the treatment of diabetes. J Nephropathol. 2016; 5(1):44-50.
- Karamati SA, Hassanzadazar H, Bahmani M, Rafieian-Kopaei M. Herbal and chemical drugs effective on malaria. Asian Pac J Trop Dis 2014; 4(Suppl 2): 599-601.
- Bahmani M, Rafieian-Kopaei M, Jeloudari M, Eftekhari Z, Delfan B, Zargaran A, Forouzan SH. A review of the health effects and uses of drugs of plant licorice (*Glycyrrhiza glabra* L.) in Iran. Asian Pac J Trop Dis 2014; 4(Suppl 2): 847-849.
- Delfan B, Bahmani M, Rafieian-Kopaei M, Delfan M, Saki K. A review study on ethnobotanical study of medicinal plants used in relief of toothache in Lorestan Province, Iran. Asian Pac J Trop Dis 2014; 4(Suppl 2): 879-884.

16. Saki K, Bahmani M, Rafieian-Kopaei M, Hassanzadazar H, Dehghan K, Bahmani F, Asadzadeh J. The most common native medicinal plants used for psychiatric and neurological disorders in Urmia city, northwest of Iran. *Asian Pac J Trop Dis* 2014; 4(Suppl 2): 895-901.
17. Bahmani M, Karamati SA, Hassanzadazar H, Forouzan SH, Rafieian-Kopaei M, Kazemi-Ghoshchi B, Asadzadeh J, Kheiri AGh, Ehsan Bahmani E. Ethnobotanic study of medicinal plants in Urmia city: identification and traditional using of antiparasites plants. *Asian Pac J Trop Dis* 2014; 4(Suppl 2): 906-910.
18. Delfan B, Bahmani M, Eftekhari Z, Jelodari M, Saki K, Mohammadi T. Effective herbs on the wound and skin disorders: a ethnobotanical study in Lorestan province, west of Iran. *Asian Pac J Trop Dis* 2014; 4(Suppl 2): 938-942.
19. Bahmani M, Rafieian M, Baradaran A, Rafieian S, Rafieian-kopaei M. Nephrotoxicity and hepatotoxicity evaluation of *Crocus sativus* stigmas in neonates of nursing mice. *J Nephropathol.* 2014; 3(2): 81-85.
20. Bahmani M, Zargaran A, Rafieian-Kopaei M. Identification of medicinal plants of Urmia for treatment of gastrointestinal disorders. *Rev Bras Farmacogn.* 2014;24(4):468-80.
21. Bahmani M, Saki K, Rafieian-Kopaei M, Karamati SA, Eftekhari Z, Jelodari M. The most common herbal medicines affecting sarcomastigophora branches: a review study. *Asian Pac J Trop Med.* 2014 Sep;7:S14-S21.
22. Sarrafchi A, Bahmani M, Shirzad H, Rafieian-Kopaei M. Oxidative stress and Parkinson's disease: New hopes in treatment with herbal antioxidants. *Curr Pharm Des.* 2016; 22(2): 238 – 246.
23. Bahmani M, Shirzad H, Majlesi M, Shahinfard N, Rafieian-Kopaei M. A review study on analgesic applications of Iranian medicinal plants. *Asian Pac J Trop Med.* 2014 Sep;7:S43-S53.
24. Sewell RDE, Rafieian-Kopaei M. The history and ups and downs of herbal medicine usage. *J Herbmed Pharmacol.* 2014; 3(1): 1-3.
25. Nasri H, Sahinfard N, Rafieian M, Rafieian S, Shirzad M, Rafieian-kopaei M. Turmeric: A spice with multifunctional medicinal properties. *J Herbmed Pharmacol.* 2014; 3(1): 5-8.
26. Shirzad H, Taji F, Rafieian-Kopaei M. Correlation between antioxidant activity of garlic extracts and WEHI-164 fibrosarcoma tumor growth in BALB/c mice. *J Med Food.* 2011; 14(9):969-74.
27. Allahghadri T, Rasooli I, Owlia P, Jalali-Nadooshan M, Ghazanfari T, Taghizadeh M, Darvish-Alipoor-Astaneh Sh. Antimicrobial Property, Antioxidant Capacity, and Cytotoxicity of Essential Oil from Cumin Produced in Iran. *Journal Of Food Science* 2010; 75(2): 54-61.
28. Ihsan Edan Alsaimary and Thuraya J. Mezaal. Evaluation of Efficiency of some Disinfectants and Antibacterial Agents on Bacterial Pathogens Isolated from Post-operative Wounds. *Advances in Bioresearch*, 2010; 1(2): 51 – 54.
29. Mohaddese Mahboubi, Ghasem Haghi. Antimicrobial activity and chemical composition of *Mentha pulegium* L. essential oil. *Journal of Ethnopharmacology* 2015; 119 (2008) 325–327.
30. Mohaddese Mahboubi, Ghasem Haghi. Antimicrobial activity and chemical composition of *Mentha pulegium* L. essential oil. *Journal of Ethnopharmacology* 119 (2008) 325–327.
31. Latif Gachkar, Davood Yadegari, Mohammad Bagher Rezaei, Masood Taghizadeh, Shakiba Alipoor Astaneh, Iraj Rasooli. Chemical and biological characteristics of *Cuminum cyminum* and *Rosmarinus officinalis* essential oils. *Food Chemistry* 102 (2007) 898–904.
32. Abdolrasoul H. Ebrahimabadi, Ebrahim H. Ebrahimabadi, Zahra Djafari-Bidgoli, Fereshteh Jookar Kashi, Asma Mazoochi, Hossein Batooli. Composition and antioxidant and antimicrobial activity of the essential oil and extracts of *Stachys inflata* Benth from Iran. *Food Chemistry* 119 (2010) 452–458.
33. S. Nejad Ebrahimi, J. Hadian, M.H. Mirjalili, A. Sonboli, M. Yousefzadi. Essential oil composition and antibacterial activity of *Thymus caramanicus* at different phenological stages. *Food Chemistry* 110 (2008) 927–931.
34. Seyyed Siavash Saei-Dehkordi, Hossein Tajik, Mehran Moradi, Farahnaz Khalighi-Sigaroodi. Chemical composition of essential oils in *Zataria multiflora* Boiss. From different parts of Iran and their radical scavenging and antimicrobial activity. *Food and Chemical Toxicology* 48 (2010) 1562–1567.
35. Peyman SALEHI, Ali SONBOLI, Fereshteh EFTEKHAR, Samad NEJAD-EBRAHIMI and Morteza YOUSEFZADI. Essential Oil Composition, Antibacterial and Antioxidant Activity of the Oil and Various Extracts of *Ziziphora clinopodioides* subsp. *rigida* (BOISS.) RECH. f. from Iran. *Biol. Pharm. Bull* 2005; 28(10) 1892—1896.
36. Mehdizadeh T and Razavi-Rouhani SM. Antibacterial activity of essential oil extracts of various onions (*Allium cepa* L.) on *Escherichia coli* and *Staphylococcus aureus*. *Journal of Agricultural Sciences and Natural Resources*; 15(2): 22-31.
37. Mohsenzadeh M. Evaluation of antibacterial activity of selected essential oils agianst of *Staphylococcus aureus* and *Eschericia coli* in Nutrient Broth Medicim. *Pakistan J Biolog Sci* 2007; 10(2): 3693-3697.
38. Maryam Abbasvali, Mahmoud Esmaeili Koutamehr, Hamdollah Moshtaghi Mohammad Hadi Eskandari. Antibacterial activity of acetonic, ethanolic and methanolic olive leaf extracts of four Iranian cultivars against *Escherichia coli*. *Journal of Food Microbiology* 2015; 2(2): 67 To 77.
39. Sharafati-chaleshtori R, Mahmoud Rafieian-kopaei M. Screening of antibacterial effect of the *Scrophularia*

- Striata against E. coli in vitro. J Herbmed Pharmacol. 2014; 3(1): 31-34.
40. Asadi SY, Parsaei P, Karimi M, Ezzati S, Zamiri A, Mohammadizadeh F, Rafieian-Kopaei M. Effect of green tea (*Camellia sinensis*) extract on healing process of surgical wounds in rat. Int J Surg. 2013;11(4):332-7.
41. Bahmani M, Vakili-Saatloo N, Gholami-Ahangaran M, Karamati SA, Khalil-Banihabib E, Hajigholizadeh Gh, et al. A comparison study on the anti-leech effects of onion (*Allium cepa* L) and ginger (*Zingiber officinale*) with levamisole and triclabendazole. J Herbmed Pharmacol. 2013; 2(1): 1-3.
42. Gupta A, Shaikh AC, Chaphalkar SR. Aqueous extract of Calamus rotang as a novel immunoadjuvant enhances both humoral and cell mediated immune response. J Herbmed Pharmacol. 2017;6(1):43-48.
43. Bahmani M, Zargaran A, Rafieian-Kopaei M. Identification of medicinal plants of Urmia for treatment of gastrointestinal disorders. Rev Bras Farmacogn 2014; 24: 468-480.
44. Baharvand-Ahmadi B, Bahmani M, Tajeddini P, Naghdi N, Rafieian-Kopaei M. An ethno-medicinal study of medicinal plants used for the treatment of diabetes. J Nephropathol. 2016; 5(1):44-50.
45. Bahmani M, Sarrafchi A, Shirzad H, Rafieian-Kopaei M. Autism: Pathophysiology and promising herbal remedies. Curr Pharm Des. 2016; 22(3):277–285. DOI: 10.2174/138161282266151112151529.
46. Baharvand-Ahmadi B, Bahmani M, Tajeddini P, Naghdi N, Rafieian-Kopaei M. An ethno-medicinal study of medicinal plants used for the treatment of diabetes. J Nephropathol. 2016; 5(1):44-50.
47. Bahmani M, Eftekhari Z, Saki K, Fazeli-Moghadam E, Jelodari M, Rafieian-Kopaei M. Obesity Phytotherapy: Review of Native Herbs Used in Traditional Medicine for Obesity. J Evid Based Complementary Altern Med. 2015 Aug 12. pii: 2156587215599105. [Epub ahead of print].
48. Bahmani M, Shirzad H, Mirhosseini M, Mesripour A, Rafieian-Kopaei M. A Review on Ethnobotanical and Therapeutic Uses of Fenugreek (*Trigonella foenum-graceum* L). J Evid Based Complementary Altern Med. 2015 Apr 27. pii: 2156587215583405. [Epub ahead of print].
49. Ebrahimie M, Bahmani M, Shirzad H, Rafieian-Kopaei M, Saki K. A Review Study on the Effect of Iranian Herbal Medicines on Opioid Withdrawal Syndrome. J Evid Based Complementary Altern Med. 2015 Oct;20(4):302-9.
50. Bahmani M, Mirhoseini M, Shirzad H, Sedighi M, Shahinfard N, Rafieian-Kopaei M. A review on promising natural agents effective on hyperlipidemia. J Evid Based Complementary Altern Med. 2015 Jul;20(3):228-38. doi: 10.1177/2156587214568457. Epub 2015 Jan 28.
51. Ebrahimi A, Chavoushpour M, Mahzoonieh MR, Lotfalian S. Antibacterial activity and ciprofloxacinpotentiation property of *Berberis vulgaris asperma* stem extracts on pathogenic bacteria. J HerbMed Pharmacol. 2016; 5(3):112-115.
52. Balkrishna A, Kumar-Manikyam H, Sharma VK, Sharma N. Clastogenic effect of *Picrorhiza kurroa* rhizome extract on cultured human peripheral blood lymphocytes. J HerbMed Pharmacol. 2016;5(4):131-136.
53. Ihekwereme CP, Agbata CA, Chukwueze KO, Agu SC. In vivo evaluation of antiplasmodial activity of hydroethanolic stem extract of *Baphia pubescens* in *Plasmodium berghei* infected albino mice. J HerbMed Pharmacol. 2016;5(4):149-152.
54. Gupta A, Chaphalkar SR. Virucidal potential of *Prosopis spicigera* and *Mangifera indica* on human peripheral blood mononuclear cells. J HerbMed Pharmacol. 2016;5(4):162-165.
55. Bahmani M, Vakili-Saatloo N, Maghsoudi R, Momtaz H, Saki K, Kazemi-Ghoshchi B, et al. A comparative study on the effect of ethanol extract of wild *Scrophularia deserti* and streptomycin on *Brucella melitensis*. J Herbmed Pharmacol. 2013; 2(1): 17-20.
56. Amirmohammadi M, Khajoenia S, Bahmani M, Rafieian-Kopaei M, Eftekhari Z, Qorbani M. In vivo evaluation of antiparasitic effects of *Artemisia abrotanum* and *Salvia officinalis* extracts on *Syphacia obvelata*, *Aspiculorhynchus tetrapetra* and *Hymenolepis nana* parasites. Asian Pac J Trop Dis. 2014; 4(Suppl 1): S250-S254.
57. Bahmani, M., Rafieian-Kopaei, M. Medicinal plants and secondary metabolites for leech control. Asian Pac J Trop Dis. 2014; 4(4): 315-316.
58. Bahmani M, Rafieian-Kopaei M, Hassanzadazar H, Saki K, Karamati SA, Delfan B. A review on most important herbal and synthetic antihelminthic drugs. Asian Pac J Trop Med 2014; 7(Suppl 1): 29-33.
59. Nasri H, Nematbakhsh M, Rafieian-Kopaei M. Ethanolic extract of garlic for attenuation of gentamicin-induced nephrotoxicity in Wistar rats. Iran J Kidney Dis. 2013; 7(5):376-82.
60. Heidarian E, Rafieian-Kopaei M. Protective effect of artichoke (*Cynara scolymus*) leaf extract against lead toxicity in rat. Pharm Biol. 2013;51(9):1104-9.
61. Rafieian-Kopaei M, Gray AM, Spencer PS, Sewell RD. Contrasting actions of acute or chronic paroxetine and fluvoxamine on morphine withdrawal-induced place conditioning. Eur J Pharmacol. 1995; 6;275(2):185-9.
62. Nasri H, Rafieian-Kopaei M. Protective effects of herbal antioxidants on diabetic kidney disease. J Res Med Sci. 2014; 19(1):82-3.
63. Baradaran A, Nasri H, Nematbakhsh M, Rafieian-Kopaei M. Antioxidant activity and preventive effect of aqueous leaf extract of *Aloe Vera* on gentamicin-induced nephrotoxicity in male Wistar rats. Clinica Terapeutica. 2014;165(1):7-11.
64. Nasri H., Rafieian-Kopaei M. Tubular kidney protection by antioxidants. Iranian J Publ Health. 2013; 42(10): 1194-1196.

65. Nasri H, Rafieian-Kopaei M. Tubular kidney protection by antioxidants. *Iranian J Publ Health*. 2013; 42(10): 1194-1196.
66. Rafieian-Kopaei M, Nasri H. Re: Erythropoietin ameliorates oxidative stress and tissue injury following renal ischemia/reperfusion in rat kidney and lung. *Med Princ Pract*. 2014;23(1):95.
67. Baradaran A, Nasri H, Rafieian-Kopaei M. Comment on: Anti-oxidative stress activity of *Stachys lavandulifolia* aqueous extract in humans. *Cell J*. 2013;15(3): 272-3.
68. Asadi-Samani M, Rafieian-Kopaei M, and Azimi N. Gundelia: A systematic review of medicinal and molecular perspective. *Pak J Biol Sci*. 2013; 16: 1238-47.
69. Nasri H, Behradmanesh S, Ahmadi A, Rafieian-Kopaei M. Impact of oral vitamin D (cholecalciferol) replacement therapy on blood pressure in type 2 diabetes patients; a randomized, double-blind, placebo controlled clinical trial. *J Nephropathol*. 2014 Jan;3(1):29-33.
70. Rafieian-Kopaei M, Nasri H, Nematbakhsh M, Baradaran A, Gheissari A, RouhiH, Ahmadi Soleimani M, Baradaran-Ghahfarokhi M, Ghaed-Amini F, Ardalan M. Erythropoietin ameliorates gentamicin-induced renal toxicity: A biochemical and histopathological study. *J Nephropathology* 2012; 1(2): 109-116.
71. Nasri H, Baradaran A, Shirzad H, Rafieian-Kopaei M. New Concepts in Nutraceuticals as Alternative for Pharmaceuticals. *Int J Prev Med* 2014;5:1487-99.
72. Ghaed F, Rafieian-Kopaei M, Nematbakhsh M , Baradaran A, Nasri H .Ameliorative effects of metformin on renal histologic and biochemical alterations of gentamicin-induced renal toxicity in Wistar rats Amini, FG. *J Res Med Sci*. 2012; 17 (7): 621-625.
73. Nasri H, Mortazavi M, Ghorbani A, Shahbazian H, Kheiri S, Baradaran A, Emami-Naieni A, Saffari M, Mardani S, Momeni A, Madihi Y, Baradaran-Ghahfarokhi M, Rafien-Kopaei M, Hedayati P, Baradaran Sh, Ardalan M, Sajjadieh Sh, Assarzadegan N and etl. Oxford-MEST classification in IgA nephropathy patients: A report from Iran. *J Nephropathol*. 2012; 1(1):31-42.
74. Rafieian-Kopaei M, Asgary S, Adelnia A, Setorki M, Khazaei M, Kazemi S, Shamsi F. The effects of cornelian cherry on atherosclerosis and atherogenic factors in hypercholesterolemic rabbits. *J Med Plants Res*. 2011; 5(13): 2670-2676.
75. Azadmehr A, Hajiaghaei R, Afshari A, Amirghofran Z, Refieian-Kopaei M, yousofi H, Darani and Hedayatollah Shirzad. Evaluation of in vivo immune response activity and in vitro anti-cancer effect by *Scrophularia megalantha*. *J Med Plants Res*. 2011; 5(11): 2365-2368.
76. Nasri H, Rafieian-Kopaei M. Tubular kidney protection by antioxidants. *Iran J Public Health*. 2013; 42(10):1194-1196.
77. Akhlaghi M, Shanian Gh, Rafieian-Koupaei M, Parvin N, Saadat M, Akhlaghi M. Citrus aurantium Blossom and Preoperative Anxiety. *Revista Brasileira de Anestesiologia* 2011; 61 (6):702-712.
78. Mirhosseini M, Baradaran A, Rafieian-Kopaei M. Anethum graveolens and hyperlipidemia: A randomized clinical trial. *J Res Med Sci* 2014;19:758-61.
79. Nasri H, Rafieian-Kopaei M. Tubular kidney protection by antioxidants. *Iran J Public Health*. 2013; 42(10):1194-1196.
80. Nasri H, Rafieian-Kopaei M. Oxidative stress and aging prevention. *Int J Prev Med*. 2013; 4(9): 1101-1102.
81. Asadi SY, Parsaei P, Karimi M, Ezzati S, Zamiri A, Mohammadizadeh F, Rafieian-Kopaei M. Effect of green tea (*Camellia sinensis*) extract on healing process of surgical wounds in rat. *Int J Surg*. 2013;11(4):332-7. doi: 10.1016/j.ijsu.2013.02.014. Epub 2013 Feb 28.
82. Parsaei P, Karimi M, Asadi SY, Rafieian-Kopaei M. Bioactive components and preventive effect of green tea (*Camellia sinensis*) extract on postlaparotomy intra-abdominal adhesion in rats. *Int J Surg*. 2013; <http://dx.doi.org/10.1016/j.ijsu.2013.08.014>.
83. Shirzad H, Shahrani M, Rafieian-Kopaei M. Comparison of morphine and tramadol effects on phagocytic activity of mice peritoneal phagocytes in vivo. *Int Immunopharmacol*. 2009 Jul;9(7-8):968-70. Epub 2009 Apr 8. PMID: 19361579.
84. Sedighi M., Rafieian-Kopaei M., Noori-Ahmabadabi M. *Kelussia odoratissima* Mozaffarian inhibits ileum contractions through voltage dependent and beta adrenergic receptors. *Life Sci J*. 9(4) 2012: 1033-1038.