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Anticancer Plants in Islamic Traditional Medicine

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Abstract

Islamic Traditional Medicine (ITM) is a holistic and comprehensive medical school that has antecedents over 12 centuries ago.

In ITM, cancer was a known disease with many options for treatment. Razi (Razes; 865-925 AD), Ahwazi (Haly Abbas; 930-994 AD), Avicenna (980-1037AD), Jorjani (1042-1136 AD), and Aqili Khorasani (18th century) are among eminent physicians who discussed different types of cancers and their management in their books. They used a large number of plant species for treatment of different tumors.

Although modern medicine has made tremendous advances in cancer control, the effectiveness of its therapeutic approach is often limited by toxic effects on other organs. Moreover, a large number of people in the world have limited or no access to cancer treatment services. Accordingly, benefiting from traditional medicine schools and effective natural medicines to prevent and control cancer would be valuable. In addition, using the teachings of such schools can lead to the discovery of new bioactive compounds and therapeutic methods.

In this chapter, a brief history of Islamic medicine and its approach to treat cancer as well as anticancer medicinal plants have been discussed. Out of 107 ITM suggested plant species, 59 plants or their chemical compounds have proven to possess cytotoxic and antitumor activities using pharmacological investigations. These findings show the profound insight of ITM physicians into cancer treatment.

Keywords: Anticancer Plants, Cancer treatment

1. Introduction

Islamic Traditional Medicine (ITM) is a holistic system of medicine which flourished during the Islamic Golden Age (750-1258 AD). It was practiced and taught throughout the Islamic territory. In that era, Muslim and non-Muslim medical scholars, especially Iranian physicians and pharmacists, translated the scientific knowledge which they inherited from ancient Greece and Iran. They endeavored to revive and develop this medical knowledge, remove superstitious ideas and faults from it, and establish an authentic medical school. Although most ITM scientists were not Arabs, the majority of their books are written in Arabic, the lingua franca of the Islamic civilization. Razi (Razhes; 865-925 AD), Ahwazi (Haly Abbas; 930-994 AD), Avicenna (980-1037AD), Jorjani (1042-1136 AD), and Aqili Khorasani (18th century) are eminent Iranian physicians who had the most contributions to ITM promotion.

Although significant progress has been made in cancer control in the last few decades, the effectiveness of modern therapeutic approach is often limited by toxic effects on other organs. Moreover, a large number of people in the world have limited or no access to cancer treatment services. Accordingly, utilizing information from traditional medicine systems to identify alternative methods to prevent and control cancer would be desirable. Furthermore, use of traditional medicine can lead to the discovery of new bioactive compounds as well as available, safe and affordable drugs.

In Islamic medicine, cancer was a known disease with many options for treatment. In the majority of ITM textbooks there is a chapter dedicated to cancer definition, symptoms, etiology, diagnosis, prevention, and management.

In this chapter we will discuss cancer, its etiology and management in the view of Islamic Traditional Medicine. In addition, ITM medicinal plants used to treat different types of cancers and modern pharmacological data confirming their traditional usage will be inserted in tables.

2. Cancer definition from ITM point of view

ITM is based on the theory of humorism which posits that the human body consists of four basic fluids, namely humors. The four humors are blood, phlegm, yellow bile, and black bile and each one corresponds to paired qualities: blood is hot and wet, phlegm is cold and wet, yellow bile is hot and dry, and black bile is cold and dry. A proper balance between humors is essential to maintaining health. Accordingly, all diseases and disabilities including cancer result from an excess or putridity of these humors.

According to ITM, cancer is a kind of black bile swelling which is accompanied by pain, pulsation, inflammation, and angiogenesis [1]. Blood vessels surrounding a tumor are overfilled and stiff and contain a dark and black blood [2]. The reason for the appellation “cancer” is due to the similarity between the shape of a cancerous tumor affecting an organ and a crab grasping its prey. It may also due to the spherical shape and darkness of tumor and origination of blood vessels from its milieu which resembles crab’s feet [3]. Cancer frequently

affects soft and porous organs and for this reason, it mainly involves breast and innervated organs (such as uterus) in females, and throat, larynx, testicles and penis in males [1].

Initially, cancer is the size of a broad bean or smaller, hard, spherical, mobile, dark, and slightly warm [3]. It will then begin to grow gradually and reach to the size of a walnut or larger. It might be curable during its early stages of development, but diagnosis is difficult in these stages. On the other hand, upon progression and appearance of clinical manifestations, treatment of cancer would be difficult or even impossible [1].

Ocular, nasal cavity, breast, uterine, liver, and other visceral organs and skin cancers are among the most frequently mentioned cancers in ITM texts. Cancers are divided into two main types: ulcerative and nonulcerative cancers.

Some cancerous tumors are easily ulcerated but some others are not. Cancerous wounds usually spread centripetally and their discharges are purulent. Use of appropriate medications can prevent the ulceration of susceptible tumors. In contrast, some cancerous tumors that are not prone to ulceration would be ulcerated following the administration of improper drugs [1].

Advanced and large tumors are very sensitive and painful, with a distinctive red to yellow color and a caustic and burning pain. Such tumors might erupt spontaneously and a purulent and bloody discharge may appear on wound surface. The resulting wounds are very sensitive and can produce corrosion in surrounding tissues [2].

3. Cancer etiology

According to ITM, excessive accumulation of abnormal black bile in a body site is the main cause of cancer. Aging, prolonged stress exposure, consuming cold and dry food items and hardwork are leading causes of increasing black bile production [1]. In some cases, hemorrhage (such as menstruation, abnormal uterine hemorrhage, or hemorrhoids bleeding) is a defending mechanism against the accumulation of bad humors in the body. Therefore, completely stopping the bleeding by surgical procedures and other medications can result in a black bile accretion and consequently increasing the risk of cancer and other diseases originating from excessive black bile (including cancer, melancholia, liver problems, psoriasis, etc.) [2].

4. Cancer management

Islamic traditional medicine suggested multiple strategies to the control and treatment of cancer. Surgical manipulation, venesection, diet adjustment, and use of natural medicines including solid, semisolid, and liquid dosage forms with oral and external route of administration are among these methods.

4.1. Surgery and manipulation

Surgery is used to eradicate tumors in their early stages of development. Small tumors which are distant from vital organs are good candidates for surgery. The tumor should be cut from

its origin and some parts of adjacent nonaffected tissues should be excised as well. In addition, bleeding should be allowed until large blood volumes come out and surrounding tissue should be pressed in order to expel blood mixed with black bile thoroughly. Afterwards, the injured site should be salved or cauterized. However, if the cancerous tumor is located in the vicinity of sensitive and vital organs, surgical procedure and cauterization would be very dangerous and may change the cancer to a nonhealing ulcer [2, 4].

Regular venesection is also suggested during the initial phases of cancer, to withdraw black bile blood from body.

4.2. Nutrition therapy

In Islamic traditional medicine, dietary recommendations have been proposed to slow the progression of advanced tumors that cannot be manipulated due to metastasis concerns. In these cases, nutritional care would increase the longevity of patients. Food items with wet temperament which produce high-quality blood like beer, almond oil, fresh small fishes, soft-boiled egg yolk, lamb, bird meat, ripe and sweet apples, sweet plums, bananas, raisin, black gram (*Vigna mungo* (L.) Hepper), spinach (*Spinacia oleracea* L.), pumpkin, light wine, and fresh cow's milk and dough are administered [1]. On the other hand, excessive intake of foodstuffs which induce black bile production in the body (such as eggplant (*Solanum melongena* L.), lentil (*Lens esculenta* Moench), date (*Phoenix dactylifera* L.), cabbage (*Brassica oleracea* L.), beef, black and thick wine, and salt-cured meat) is severely restricted [5].

4.3. Pharmacotherapy

As mentioned above, from the Islamic Traditional Medicine point of view, an excess of abnormal black bile in various body sites can lead to cancer formation. In order to treat cancer, black bile should be expelled from the body by using an appropriate purgative, and then preventing the generation and accumulation of black bile in vessels as far as possible [2]. For this purpose, many single and compound medications of herbal, animal, and mineral origin have been recommended.

The essential point in pharmacotherapy of cancers and tumors is avoiding the use of caustic and irritant medications to prevent further stimulation and ulceration [2].

Cancer medications can be administered internally (oral, enemas, vaginal douches, etc.) or applied topically (cataplasms, topical oils, liniments, lotions, dusting powders, etc.) [1, 3]. Administration of local anticancer drugs has the following purposes: cancer eradication, metastasis prevention, preventing ulceration, and healing ulcerated tumors.

5. Mechanisms of action of anticancer drugs

Anticancer drugs act through different mechanisms:

- i. **Black bile purgatives** are common anticancer drugs which can facilitate elimination of abnormal black bile from whole body. Purgatives should be administered fre-

quently. From the ITM point of view, clover dodder (*Cuscuta epithimum* Murr.) is the most valuable black bile purgative, which is commonly used to treat diseases caused by a surplus or imbalance of this humor such as all types of cancer, melancholia, leprosy, vitiligo, etc. For this purpose, a mixture of clover dodder with cheese whey or hydromel should be administered frequently. A decoction of the plant in oxymel is also prescribed [3]. Common polypody (*Polypodium vulgare* L.), French lavender (*Lavandula stoechas* L.), colocynth [*Citrullus colocynthis* (L.) Schrad.], and black hellebore (*Helleborus niger* L.) are other potent black bile purgatives.

- ii. **Antiulcer drugs** can inhibit ulceration of tumors. *Urtica pilulifera* L. and *Aloe vera* (L.) Burm.f. are examples of such plants.
- iii. **Wound-healing medications** accelerate healing of cancerous wounds. *Althaea officinalis* L., *Brassica oleracea* L., and *Viola odorata* L. have wound-healing activity.
- iv. **Analgesic drugs** relieve cancer pain. *Parietaria officinalis* L. and *Solanum nigrum* L. are plants with analgesic effect.

6. Anticancer plants

One hundred and seven plant species from 53 families have been mentioned to be effective in the management and curing of different types of cancers. Plants Latin and Arabic names, their families, medicinal parts, dosage forms, and routes of administration are given in Table 1.

Scientific name	Family	Arabic common name	Part used	Cancer type	Administration and locality	Ref.
<i>Acacia nilotica</i> (L.) Delile syn. <i>Acacia arabica</i> (Lam.) Willd.	Fabaceae	samgh	G	eye	ocular suppository	[3]
<i>Allium ampeloprasum</i> L. syn. <i>Allium porrum</i> L.	Liliaceae	korrath	Le	all types	decoction(O)	[6]
<i>Aloe vera</i> (L.) Burm.f.	Liliaceae	sabr	Sa	uterus ulcerating tumor internal organs	unguent powder(E) decoction(O) pill(O)	[1, 3]
<i>Alpinia officinarum</i> Hance	Zingiberaceae	khawlanjan	Rh	all types	electuary(O)	[8, 9]
<i>Althaea officinalis</i> L.	Malvaceae	khatmi	R	all types uterus	liniment cataplasm vaginal bath	[1, 3] [2, 7]
<i>Alyssum homalocarpum</i> (Fisch. & C. A.Mey.) Boiss.	Brassicaceae	tudari urisimun	Se	internal organs skin	cataplasm	[2, 5, 9-11]

Scientific name	Family	Arabic common name	Part used	Cancer type	Administration and locality	Ref.
<i>Amaranthus blitum</i> L.	Amaranthaceae	baghlat al-yamaaniah	Ap	uterus	cataplasm	[7]
<i>Anchusa azurea</i> Mill. syn. <i>Anchusa italica</i> Retz.	Boraginaceae	lesan al-thawr	Le	all types	syrup	[7]
<i>Anethum graveolens</i> L.	Apiaceae	shebeth	Ap	uterus	unguent topical oil	[1, 3]
<i>Beta vulgaris</i> L.	Chenopodiaceae	salq	Le	all types	decoction(O)	[6]
<i>Boswellia sacra</i> Flueck. syn. <i>Boswellia carteri</i> Birdw.	Burseraceae	kondor	Ogr	eye	condensed smoke as collyrium	[2, 3, 5, 8, 9]
<i>Brassica oleracea</i> L.	Brassicaceae	karnab	Le Fl	uterus all types skin	unguent decoction(E) vaginal bath cooked(O) cooked (E)	[1, 3, 7] [6, 8, 9]
<i>Capparis spinosa</i> L.	Capparaceae	kabar	Rb	uterus all types	unguent cataplasm	[3] [1, 5]
<i>Carthamus tinctorius</i> L.	Asteraceae	moasfar	Se	all types	raw seeds(O)	[6]
<i>Carum carvi</i> L.	Apiaceae	kerawia	Se	all types	raw seeds(O)	[6]
<i>Cassia fistula</i> L.	Fabaceae	khiair shanbar	Lg	tongue ulcerated cancer uterine	extract (E)	[3] [12]
<i>Chamaemelum nobile</i> (L.) All.syn. <i>Anthemis nobilis</i> L.	Asteraceae	babunaj	Fl	uterus	unguent topical oil decoction(E)	[1, 3, 7, 12]
<i>Cicer arietinum</i> L.	Fabaceae	homs	Se	skin all types ulcerating tumor	cooked(O) infusion(E) cataplasm	[8, 9] [2]
<i>Cichorium intybus</i> L.	Asteraceae	hindeba	Le	uterus all types ulcerating tumor	extract(E) cataplasm extract(O) cataplasm	[1, 6, 8] [7]
<i>Cinnamomum cassia</i> (L.) J.Presl	Luraceae	salikhah	Br	liver	electuary	[2]
<i>Cistus creticus</i> L. syn. <i>Cistus ladaniferus</i> Curtis	Cistaceae	ladan	Fl	all types	oil (E)	[3]

Scientific name	Family	Arabic common name	Part used	Cancer type	Administration and locality	Ref.
<i>Citrullus colocynthis</i> (L.) Schrad.	Cucurbitaceae	hanzal	Fr	internal organs	decoction(O) pill(O)	[1]
<i>Commiphora mukul</i> (Hook. ex Stocks) Engl.	Burseraceae	moql	Ogr	uterus	unguent	[1, 3, 7]
<i>Convolvulus pseudoscammonia</i> C. Koch	Convolvulaceae	saqmunia	Rdj	internal organs	decoction(O) pill(O)	[1]
<i>Cordia myxa</i> L.	Boraginaceae	debgh	Fr	uterine cancer	decoction(O)	[12]
<i>Coriandrum sativum</i> L.	Apiaceae	kozborah	Le Se	eye	extract(E)	[1]
				uterus	cataplasm	[1, 3]
				skin	vaginal douche	[2, 7, 9]
				all types	cataplasm	[8, 9]
					extract (E)	[6, 7]
<i>Crocus sativus</i> L.	Iridaceae	za`afaran	St	eye	cataplasm	[3]
				uterus	cataplasm	[1, 2, 11,
				metastatic	cataplasm	13]
				ulcerating tumor	cataplasm	[12]
				liver	electuary	[2]
<i>Cucurbita maxima</i> Duchartre, <i>Cucurbita pepo</i> L.	Cucurbitaceae	qar`a	Fr	internal organs	cooked (O) extract lotion	[1, 6-8]
<i>Cuscuta epithymum</i> (L.) L.	Cuscutaceae	aftimun	Ap	all types ulcerating tumor	powder(O) decoction(O)	[1-4, 6, 8-10]
<i>Cymbopogon schoenanthus</i> (L.) Spreng.	Poaceae	ezkher	Fl	liver	electuary	[2]
<i>Cynara scolymus</i> L.	Asteraceae	kankar zad	G		cataplasm	[2]
<i>Dorema ammoniacum</i> D. Don	Apiaceae	oshaq	Ogr	all types	unguent	[3, 8, 14]
<i>Dracunculus vulgaris</i> Schottsyn. <i>Arum</i> <i>dracunculus</i> L.	Araceae	luf al-hayyah	Be Se	nose breast testicle	extract decoction(E)	[2, 3, 5, 8-10] [2]
<i>Ecballium elaterium</i> (L.) A. Rich.	Cucurbitaceae	qetha al-hemar	Fr	all types	liniment(E)	[1, 3]
<i>Erysimum</i> × <i>cheiri</i> (L.) Crantz syn. <i>Cheiranthus</i> × <i>cheiri</i> L.	Brassicaceae	kheiri	Fl	all types uterus	oil(E)	[3, 7] [12]
<i>Ferula gummosa</i> Boiss.	Apiaceae	ghennah	Ogr	all types	unguent	[8]
<i>Ficus carica</i> L.	Moraceae	teen	Fr	tongue	cooked(E)	[3]
				all types	cooked (O)	[6]

Scientific name	Family	Arabic common name	Part used	Cancer type	Administration and locality	Ref.
				ulcerating tumor uterus	cataplasm cataplasm	[12, 13] [7]
<i>Glycyrrhiza glabra</i> L.	Fabaceae	sous	R	all types uterus	syrup vaginal douche	[7, 15]
<i>Helleborus niger</i> L.	Ranunculaceae	kharbaq aswad	R	internal organs	decoction(O) pill(O)	[1, 7, 10]
<i>Hordeum vulgare</i> L.	Poaceae	sha`eer	Se	all types	decoction(O)	[1, 3, 6, 7]
<i>Inula helenium</i> L.	Asteraceae	rasan	R	skin	lotion	[14]
<i>Iris × germanica</i> L. syn. <i>Iris × florentina</i> L.		irsa	R	uterus	unguent	[3]
<i>Jasminum sambac</i> (L.) Aiton <i>Aegle marmelos</i> (L.) Corrêa ex Roxb.		razeqi	Fl	uterus	unguent topical oil	[3]
<i>Juglans regia</i> L.	Juglandaceae	Jawz	G	ulcerating tumor	dusting powder(E)	[2, 14]
<i>Lactuca sativa</i> (L.) Mill.	Asteraceae	khas	Le	all types uterus	extract(E) cataplasm	[1, 3] [6, 7]
<i>Laurus nobilis</i> L.	Lauraceae	hab al-ghar	Se	tongue uterus	cataplasm cataplasm	[1, 3] [7]
<i>Lavandula stoechas</i> L.	Lamiaceae	ostokhoddu	Ap	internal organs	decoction(O) pill(O)	[1]
<i>Lawsonia inermis</i> L.	Lythraceae	henna	Le	uterus	unguent oil(E)	[3]
<i>Lens culinaris</i> Medik. syn. <i>Lens esculenta</i> Moench	Fabaceae	adas	Se	uterus ulcerating tumor	vaginal douche cataplasm	[3] [7]
<i>Lilium candidum</i> L.	Liliaceae	susan	Fl	uterus	unguent oil (E)	[1, 3]
<i>Linum usitatissimum</i> L.	Linaceae	katan	Se Mu	all types uterus	unguent oil (E) vaginal bath decoction (E)	[3] [1, 2, 7]
<i>Liquidambar orientalis</i> Mill.	Altingiaceae	mi`a	Ba	all types	unguent	[3]
<i>Lycium afrum</i> L.	Solanaceae	hozoz	Le Fs	uterus	extract vaginal douche	[15]
<i>Malva pusilla</i> Sm. syn. <i>Malva rotundifolia</i> L.	Malvaceae	khabaazi	Fl	uterus	vaginal bath cataplasm	[7]
<i>Melilotus officinalis</i> (L.) Pall.	Fabaceae	eklil al-malek	pod	eye	cataplasm	[1, 3]

Scientific name	Family	Arabic common name	Part used	Cancer type	Administration and locality	Ref.
				uterus		[2]
<i>Melissa officinalis</i> L.	Lamiaceae	badranjbuyeh	Le	all types	syrup	[7]
<i>Myrtus communis</i> L.	Myrtaceae	as	Se Le	all types	topical oil extract enema	[14]
<i>Narcissus tazetta</i> L.	Amaryllidaceae	narjes	Fl	uterus	unguent topical oil	[1, 3]
<i>Olea europaea</i> L.	Oleaceae	zeytun	Fr	uterus metastatic cancerous wounds	topical oil oil sediment (E) cataplasm	[1, 3] [13]
<i>Opopanax chironium</i> W.D.J.Koch	Apiaceae	jawshir	Ogr	all types	unguent	[3, 14]
<i>Origanum syriacum</i> L. syn. <i>Origanum maru</i> L.	Lamiaceae	mormahur	Ap	metastatic cancerous wounds	cataplasm	[13]
<i>Papaver somniferum</i> L. var. <i>album</i> (Mill.) M.A. Veselovskaya	Papaveraceae	khashkhash	Se Sa	eye uterus	ocular suppository cataplasm	[3] [1-3, 5, 12]
<i>Parietaria officinalis</i> L.	Urticaceae	hashishatah al- zojaj	Le	all types	extract(E)	[5]
<i>Phoenix dactylifera</i> L.	Arecaceae	tamr	Fr	tongue	decoction(E)	[3]
<i>Physalis alkekengi</i> L.	Solanaceae	kaknaj	Fr	all types uterus	extract(E) unguent	[7] [15]
<i>Pinus gerardiana</i> Wall. ex D.Don	Pinaceae	jalghuzah	N	all types		[6]
<i>Pinus</i> spp.	Pinaceae	ratinaj	Re	uterus	cataplasm	[7]
<i>Pistacia atlantica</i> Desf. <i>Pistacia terebinthus</i> L.	Anacardiaceae	elk al-anbat	Ogr	uterus all types	unguent	[1, 3, 6, 8]
<i>Pistacia lentiscus</i> L.	Anacardiaceae	mastaki	Ogr	all types	unguent oil(E)	[1, 3] [6]
<i>Plantago indica</i> L. syn. <i>Plantago psyllium</i> L.	Plantaginaceae	bazr qotuna	Se Mu	ulcerating tumor intestine all types	liniment enema	[3, 8] [14]
<i>Plantago major</i> L.	Plantaginaceae	lesan al-hamal	Se Mu Le	uterus ulcerating tumor	vaginal douche cataplasm	[1-3, 7, 12]
<i>Plantago ovata</i> Forssk.	Plantaginaceae	isbaghul	Se Mu	all types uterus	vaginal douche	[1, 6] [7]

Scientific name	Family	Arabic common name	Part used	Cancer type	Administration and locality	Ref.
<i>syn. Plantago ispaghula</i> Roxb. ex Fleming						
<i>Platanus orientalis</i> L.	Plantanaceae	dolb	Le	skin	decoction (E) vaginal bath	[2]
<i>Polygonum aviculare</i> L.	Polygonaceae	asa al-ra`ee	Ap	uterus	unguent extract (E)	[7, 15]
<i>Polypodium vulgare</i> L.	Polypodiaceae	basfayaj	Rh	internal organs	decoction(O) pill(O)	[1, 7]
<i>Polyporus officinalis</i> (Vill.) Fr.	Polyporaceae	ghariqun		all types	decoction(O)	[4, 7]
<i>Portulaca oleracea</i> L.	Portulacaceae	rejlah	Le	ulcerating tumor all types uterus	extract liniment enema extract(O)	[3, 8] [1] [6] [7]
<i>Prunus dulcis</i> (Mill.) D.A.Webb	Rosaceae	samgh al-llawz	G So Se	uterus all types	unguent oil (O)	[1, 3, 6, 12]
<i>Punica granatum</i> L.	Punicaceae	romman	Fr Fp	nose ulcerating tumor	juice(O)	[1] [7]
<i>Raphanus raphanistrum</i> subsp. <i>sativus</i> (L.) Domin <i>syn. Raphanus sativus</i> L.	Brassicaceae	fojl	tro	all types	decoction(O)	[6]
<i>Rhus coriaria</i> L.	Anacardiaceae	sumaagh				
<i>Ricinus communis</i> L.	Euphorbiaceae	kherwa`a	Se	uterus	unguent oil (E)	[1, 3]
<i>Rosa × damascena</i> Herrm.	Rosaceae	ward	Fl	eye uterus ulcerating tumor	oil(E) cataplasm vaginal douche oil	[1] [2, 3, 7, 12]
<i>Salix aegyptiaca</i> L.	Salicaceae	khelaf	W	skin	dusting powder(E)	[2]
<i>Sempervivum tectorum</i> L.	Crassulaceae	hay al-alam	Fl	all types	extract liniment	[3, 7]
<i>Sesbania bispinosa</i> (Jacq.) W. Wight	Fabaceae	sisban	Le		cataplasm	[9]
<i>Sisymbrium officinale</i> (L.) Scop. <i>syn. Erysimum officinale</i> L.	Brassicaceae	arismun, khabbe	Se	hard swelling cancer nonulcerating tumor	cataplasm cataplasm cataplasm cataplasm	[8, 9] [2]

Scientific name	Family	Arabic common name	Part used	Cancer type	Administration and locality	Ref.
				ear breast testicle		
<i>Smilax china</i> L.	Smilacaceae	khashab sini	R	skin	decoction(O)	[8]
<i>Solanum americanum</i> Mill. syn. <i>Solanum nigrum</i> L.	Solanaceae	enab al-tha'alab	Le	uterus all types skin	cataplasm extract cataplasm	[1-3, 6, 7] [7] [5, 8-10]
<i>Spinacia oleracea</i> L.	Chenopodiaceae	esfanakh	Le	internal organs	cooked (O)	[1, 6]
<i>Tamarix gallica</i> L.	Tamaricaceae	tarfa	Le R Sh	all types spleen	decoction(O)	[5]
<i>Tanacetum parthenium</i> (L.) Sch.Bip.	Asteraceae	oqhowan	Fl	skin uterus	extract cataplasm unguent oil(E)	[9] [1, 3]
<i>Terminalia bellirica</i> (Gaertn.) Roxb.	Combretaceae	amolaj	Fr	all types	liniment	[7]
<i>Terminalia chebula</i> Retz.	Combretaceae	ehalilaj kaboli	Fr	internal organs	decoction(O) pill(O)	[1]
<i>Trigonella foenum-graecum</i> L.	Fabaceae	holbah	Se mu	tongue uterus all types	infusion(E) oil(E) decoction(E)	[3] [1, 6, 7]
<i>Triticum spelta</i> L.	Poaceae	khondorus	Se	all types	baked(O)	[6]
<i>Urtica pilulifera</i> L.	Urticaceae	anjorah	Se Le	all types skin ulcerating tumor	cataplasm ash dusting powder(E)	[3, 8, 9, 11, 13] [2]
<i>Valeriana celtica</i> L.	Valerianaceae	sonbol	R	liver	electuary	[2]
<i>Vigna mungo</i> (L.) Heppersyn. <i>Phaseolus mungo</i> L.	Fabaceae	maash	Se	internal organs	cooked(O)	[1, 6]
<i>Viola odorata</i> L.	Violaceae	banafsaj	Ap	uterus	decoction(E) vaginal bath	[7, 15]
<i>Vitis vinifera</i> L.	Vitaceae	enab	Urf verjuice Rf Fr Fl	all types tongue all types skin	cataplasm concentrated juice(O) raw Fr(O) cataplasm	[3, 6, 7] [6] [6, 10] [10] [12]

Scientific name	Family	Arabic common name	Part used	Cancer type	Administration and locality	Ref.
				ulcerating tumor uterine cancer uterus	decoction(E) unguent	
<i>Zingiber officinale</i> Roscoe	Zingiberaceae	zanjabil	Rh	liver skin	electuary lotion	[2] [14]
<i>Ziziphus jujuba</i> Mill. syn. <i>Ziziphus vulgaris</i> Lam.	Rhamnaceae	onnab	Fr	ulcerating tumor uterus	decoction(O)	[12]

Use: E = external use, O = oral use. Part used Ap = aerial parts, Ba = balsam, Be = berries, Br = bark, Ff = fresh fruit, Fl = flowers, Fp = fruit pericarp, Fr = fruit, G = gum, Gre = gum resin, Le = leaves, Mu: mucilage, N = nuts, O = oil, Ogr = oleo-gum resin, Ore = oleoresin, R = root, Rb = root bark, Re = resin, Rdj = Root dry juice, Rf = Ripe fruit, Rh = rhizome, Se = seeds, Sh =shoots, So = seed oil, St = stigma, Tap root = Tro, Sa = sap, Urf = Unripe fruit, W = wood, Wp = whole plant.

Table 1. Medicinal plants mentioned in Islamic Traditional Medicine textbooks effective in the management of cancer.

7. Anticancer and cytotoxic activities of ITM plant species

Many ITM-suggested plants have been shown to exert anticancer activities with different mechanisms such as cytotoxic, antitumor, antiproliferative, cytostatic, and cell-migration-inhibiting effects. Pharmacological data reinterpreting ITM evidence of cancer phytotherapy are given in Table 2.

Species	Plant part(s)/compound	Solvent	Activity	Cell line	Ref.
	Stem bark	ethanol		K-562, Raji, Jurkat, HEL, Colo38, HL-60, CEM, B-16, MCF-7 and MDA-MB-231	[16-18]
		methanol, ethyl acetate and n-hexane	cytotoxic	Brine shrimp	[19]
<i>Aegle marmelos</i> (L.) Corrêa ex Roxb.	Stem bark, roots and leaves/Zeorin, dustanin, aegeline, epilupeol, lupenone and marmin	-		CEM-SS	[20]
	1-hydroxy-5,7-dimethoxy-2-naphthalene-carboxaldehyde (marmelin)	-	antitumor	HCT-116 colon cancer cell tumor xenograft in nude mice	[21]
<i>Allium porrum</i> L.	Bulbs/12-keto-porrigenin and 2,3-seco-porrigenin	-	antiproliferative	J774, WEHI 164, P388 and K3R-1	[22]

Species	Plant part(s)/compound	Solvent	Activity	Cell line	Ref.
	Bulbs/Porrigenins A and B	-		IGR-1, J774, WEHI 164 and P388	[23]
	Saponins	-		J774 and WEHI 164	[24]
<i>Aloe vera</i> (L.) Burm.f.	Leaves	aqueous ethanolic extract (70%)	cytotoxic	K562	[25]
	Flavonoids	-		MCF-7	[26]
	Di(2-ethylhexyl)phthalate	-		K562, HL60 and U937	[27]
				Neuroblastoma cells (IMR-32, IMR-5, AF8, and SJ-N-KP), pPNET cells (TC32) and Ewing's sarcoma cells (TC106)	[28]
	Aloe-emodin	-		PC3	[29]
				MCC	[30]
				U-373MG and U87 glioma cells	[31, 32]
				T24 human bladder cancer cells	[33]
				Gastric cancer cells	[34]
				MCF-7	[35]
			antitumor	Ehrlich ascitis carcinoma (EAC)	[36]
<i>Anthemis nobilis</i> L.	Sesquiterpene lactones	-	cytotoxic	HeLa and KB	[37]
<i>Beta vulgaris</i> L.	Root extract and betanin	water	cytotoxic and antitumor	MCF-7, PC3 and HepG2	[38-41]
<i>Boswellia carteri</i> Birdw.	α and β -boswellic acid acetate from resin	-	cytostatic-inhibits cell migration	KB, HCT-8, A2780 and B16F10	[42]
	Triterpene acids from resin	-	cytotoxic	HT-1080	
	Frankincense oil			IMR-32, NB-39 and SK-N-SH	[43]
	Verticilla-4(20),7,11-triene			The bladder carcinoma J82	[44]
<i>Brassica oleracea</i> L.	Sulforaphane	-		MDAH 2774 and SkOV-3	[46]
	Erucin	-		CACO-2, HL60, K562 and LNCaP	[47]
	Leaves	acidified methanol	cytotoxic	HeLa and HepG2	[48]
	2-Pyrrolidinone reach extract			PC3 and HeLa	[49]
	Brassinin	-		PC3	[50]

Species	Plant part(s)/compound	Solvent	Activity	Cell line	Ref.
<i>Capparis spinosa</i> L.	Polysaccharides and alkaloids from fruits	water	cytotoxic and antitumor	HepG2 and transplanted ascites tumor H22	[51, 52]
<i>Carthamus tinctorius</i> L.	<i>Carthamus tinctorius</i> and <i>Panax ginseng</i>	-	cytotoxic and antitumor	MDA-MB-231 breast cancer cell and normal human mammary gland cell lines	[53]
	Polysaccharide	-	antitumor	T739 lung cancer and S180 Sarcoma in mice	[54]
	Not mentioned	methanol	cytotoxic	HT-29	[55]
<i>Carum carvi</i> L.	Thymoquinone from seeds	-	cytotoxic	TNBC cells	[56]
	Rhein from flowers	-		COLO 320 DM	[57]
	Stem bark	methanol			[58]
<i>Cassia fistula</i> L.	Furanoflavones from stem bark	-	cytotoxic	NB4, A549, SHSY5Y, PC3, and MCF7	[59]
	Fistulaquinone A from fruits	-		NB4 and PC3	[60]
<i>Cicer arietinum</i> L.	C-25 protein	-	cytotoxic	KB cell line	[61]
	Leaves	chloroform	antiproliferative	HeLa, MCF7 and A431	[62]
<i>Cichorium intybus</i> L.	Lactucin and 13,14-secostigma 9(11),14(15)-dien-3 α -ol	-	cytotoxic	A2780	[63]
	Cinnamaldehyde	-		HL60	[64]
	Bark	methanol/hexane/ ethyl acetate		HeLa, A549, SK-OV-3, SK-MEL-2, XF-498 and HCT-15	[65]
<i>Cinnamomum cassia</i> (L.) J.Presl	2'-	aqueous	cytotoxic	SiHa	[66]
	Benzoyloxycinnamaldehyde	-		LNCaP, PC-3 and DU145	[67]
	Coumacasia	-		HL60 and A549	[68]
<i>Cistus creticus</i> L.				KB, P-388; and NSCLC-N6	[69]
	Labdane-type diterpenoids from leaves	-	cytotoxic	CCRF-CEM, MOLT3, H33AJ-JA-13, HUT78, H9, KM3, NAMALWA, JIYOYE, DAUDI, SDK, K562, HL60 and U973	[70-72]
	Shoot	ethanol		HeLa, MDA-MB-453 and FemX	[73]
<i>Commiphora mukul</i> Engl.	Guggulsterones	-	cytotoxic	PC3, HL60 and U973	[74]
<i>Coriandrum sativum</i> L.	Linalool	-	antitumor	Sarcoma-180 solid tumor	[75]
	Leaves	ethanol	cytotoxic	HT-29	[76]

Species	Plant part(s)/compound	Solvent	Activity	Cell line	Ref.
<i>Crocus sativus</i> L.	Stigma	ethanol	antitumor	Sarcoma-180 (S-180), Ehrlich ascites Carcinoma (EAC) and Dalton's lymphoma ascites (DLA)	[77]
				HeLa, A549 and HepG2	[78, 79]
	Crocin, crocetin, safranal and picrocrocin	-	cytotoxic	HeLa, MCF-7, PC3, k562 and HepG2	[80-86]
	Crocin		antitumor	C26 colon carcinoma	[87]
<i>Cucurbita maxima</i> Duchartre,	Seeds	methanol	cytotoxic	Brine shrimp	[88]
	Aerial parts		antitumor	Ehrlich ascites Carcinoma	[89]
	Triterpenes	-	cytotoxic	HL60 and P388	[90]
<i>Cucurbita pepo</i> L.	Leaves	hydroalcoholic	cytotoxic	HepG2 and CT26	[91]
<i>Cuscuta epithymum</i> Mur.	Aerial parts	chloroform and hydroalcoholic	cytotoxic	HeLa, HT29 and MDA-MB-46	[92]
<i>Ecballium elaterium</i> (L.) A. Rich.	Cucurbitacin E	-	cytotoxic	ZR-75-1, COLO 679, OV_95_CC3 and PC-3	[93, 94]
	Fruits	water		AGS and KYSE30	[95]
	Cucurbitacin-D	-		NSCLC-N6	[96]
<i>Ficus carica</i> L.	Leaves, fruits and latex	ethanol, ethyl acetate and dichloromethane	cytotoxic	HeLa	[97]
<i>Glycyrrhiza glabra</i> L.	Root		cytotoxic	4T1	[98]
<i>Inula helenium</i> L.	Root	methanol	cytotoxic	HT-29, MCF-7, Capan-2 and G1	[99]
		ethanol	cytotoxic	MDA-MB-23	[100]
<i>Jasminum sambac</i> (L.) Aiton	Leaves	ethanol	cytotoxic	Brine shrimp	[101]
<i>Juglans regia</i> L.	Juglanin A and B	-		Hep G2	[102]
	Leaves	chloroform		BHY, MCF7, and HT-29	[103]
	Juglanones A and B	-	cytotoxic	A549, MCF-7, BEL-7402, HeLa, COLO205, BGC-823, and SK-OV-3	[104]
	5,7-Dihydroxy-3,4'-dimethoxyflavone and regiolone			MCF-7 and BHY	[105]
<i>Lactuca sativa</i> (L.) Mill.	Sesquiterpene lactones	-	cytotoxic	HeLa and HCT-116	[106]
<i>Laurus nobilis</i> L.	Leaves	n-hexane	cytotoxic	Brine shrimp	[107]
	Sesquiterpene lactones	-		Jurkat	[108]

Species	Plant part(s)/compound	Solvent	Activity	Cell line	Ref.
				, HL-60 and LoVo	
				A2780	[109]
	Essential oil	-		C32, ACHN, LNCaP, and MCF-7	[110]
<i>Lawsonia inermis</i> L.	Bicoumarin, biflavonoid, and biquinone compounds from flowers	-	cytotoxic	MCF-7, Hela, HCT-116, and HT-29	[111]
<i>Linum usitatissimum</i> L.	Secoisolariciresinol and anhydrosecoisolariciresinol	-	cytotoxic	MCF-7 and MDA-MB-231	[112]
	Essential oil	-	cytotoxic	A549, MCF-7, Caco-2, HL-60, K562 and B16F10	[113]
<i>Melissa officinalis</i> L.	Aerial parts	ethanol		HCT-116	[114]
	Leaves	water	cytotoxic and antitumor	MCF-7, MDA-MB-468 and MDA-MB-231; DMBA-induced mammary tumors	[115]
	Citral and essential oil	-	cytotoxic	GBM	[116]
<i>Myrtus communis</i> L.	Phloroglucinols	-	cytotoxic	MT-4 cells, HepG2 and DU145	[117]
	Alkaloids	-			[118]
<i>Narcissus tazetta</i> L.	Stems and leaves	alkaloid extraction	cytotoxic	HL-60, K562, KT1/A3, and A3	[119]
	Erythrodiol	-		HL-60	[120]
<i>Olea europaea</i> L.	Hydroxytyrosol rich extract from leaves	methanol/water	cytotoxic	MCF-7	[121]
	Maslinic acid	-		HT29	[122]
	Tyrosol esters	-		MCF10A	[123]
<i>Papaver somniferum</i> L. var. <i>album</i> (Mill.) M.A.	Noscapine	-	anticancer	Refractory Multiple Myeloma, Non-Hodgkin's Lymphoma and Chronic Lymphocytic Leukemia	[124]
<i>Physalis alkekengi</i> L.	5 α -Hydroxy-25,27-dihydro-4,7-didehydro-7-deoxyneophysalin A	-	cytotoxic	PC-3 and LNCaP	[125]
	Physalin	-		HT1080 and A375-S2	[126, 127]
<i>Pistacia atlantica</i> Desf.	Polyphenol-rich extract	ethanol/water	cytotoxic	HT29	[128]
<i>Pistacia lentiscus</i> L.	Gum extract	hexane	cytotoxic	HCT116	[129]
<i>Plantago major</i> L.	Leaves	hot water	cytotoxic	Antileukemia and anticarcinoma	[130]

Species	Plant part(s)/compound	Solvent	Activity	Cell line	Ref.
	Luteolin-7-O- β -glucoside	-		TK-10, MCF-7 and UACC-62	[131]
<i>Platanus orientalis</i> L.	Flavonoids	-	cytotoxic	Human leukemic cell lines and skin cancer cell lines	[132-136]
<i>Polygonum aviculare</i> L.	Aerial parts	methanol	cytotoxic	MCF-7	[137]
<i>Portulaca oleracea</i> L.	Triterpenoids	-		HepG2	[138]
	Not mentioned	methanol		CNE-1, HeLa and HT-29 and MCF-7	[139]
	Seed		cytotoxic	HepG2	[140]
	Alkaloids	-		K562, A549, MCF-7 and MDA-MB-435	[141]
<i>Punica granatum</i> L.	Genistein and extract	not mentioned	cytotoxic	MCF-7, MMTV-Wnt-1,	[142]
	Polyphenols	juice	cytotoxic and antitumor	BT-474 and MDA-MB-231	[143]
	Fruit rind extract and fowers	methanol	cytotoxic	A549 and MCF-7	[144-146]
	Galactomannan polysaccharide	-	cytotoxic and antitumor	A375, HCT116, and HepG2; DLA and EAC murine ascites and EAC solid tumor mouse models	[147]
	4-(Methylthio)-3-butenyl isothiocyanate	-	cytotoxic	murine leukaemia cell line (L1210)	[148]
<i>Raphanus sativus</i> L.	4-Methylthio-butanyl derivatives	-	cytotoxic	A549, SK-OV-3, SK-MEL-2, and HCT-15	[149]
<i>Ricinus communis</i> L.	Leaves	volatile oil	cytotoxic	SK-MEL-28 and HeLa	[150]
<i>Rosa \times damascena</i> Mill.	Flowers	volatile oil	cytotoxic	SW742	[151]
<i>Smilax china</i> L.	Phenylpropanoid glycosides			KB, HeLa, DLD-1, MCF-7, A-549 and Med	[152]
	Kaempferol-7-O-beta-D-glucoside	-	cytotoxic	A375 and HL60	[153]
	Polyphenols			MCF-7 and MDA-MB-231	[154]
<i>Solanum nigrum</i> L.	Steroidal glycosides	-	cytotoxic	HT-29, HCT-15, LNCaP, PC-3, T47D, HepG2, NCI-H460, MCF-7, SF-268 and MDA-MB-231	[155, 156]
	Glycoprotein			MCF-7, HCT-116 and HT-29	[157-160]
	Aerial parts	methanol		HeLa and Vero	[161]
	Leaves	water	cytotoxic	AU565	[162]
	Not mentioned	hydro-alcoholic		HepG2 and CT26	[91]

Species	Plant part(s)/compound	Solvent	Activity	Cell line	Ref.
	Solamargine	-		K562	[163]
	Polyphenol rich extract	water		PZ-HPV-7	[164]
	Berries	ethanol		Jurkat and HL-60	[165]
<i>Tanacetum parthenium</i> L.	Parthenolide	-	anticancer	Leukemia	[166]
<i>Terminalia chebula</i> Willd. ex Flem.	Tannins	-	cytotoxic	A-549, SK-OV-3, SK-MEL-2, XF-498 and HCT-15	[167]
	Fruits	methanol		HOS-1	[168]
<i>Trigonella foenum-graecum</i> L.	Seeds	water	cytotoxic	HL-60 TCP, B-cell lymphomas, FRO and MCF7	[169]
	Diosgenin	-		A549	
<i>Viola odorata</i> L.	Cyclotides	-	cytotoxic	MCF-7 and MCF-7/ADR	[170]
	Resveratrol	-		3T6 and HL60	[171]
<i>Vitis vinifera</i> L.	Seed extract	methanol	cytotoxic	KB cells	[172]
	Viniferin-enriched extracts	ethanol/water		HCC1954, HCC1500 and MCF7	[173, 174]
<i>Zingiber officinale</i> Roscoe	[6]-Paradol and structurally related compounds	-	cytotoxic	KB	[175]
	Gingerols and diarylheptanoids	-		HL-60, A431, K562, HeLa, HCT-116, HT-29 and K562/ADR	[176-181]

Table 2. Anticancer activities of ITM plant species.

8. Conclusion

Taken together, it can be concluded that what physicians of Islamic medicine used for cancer treatment is proven through modern research. Out of 107 plant species which are introduced in Islamic Iranian medicine for cancer treatment, 59 plants or their chemical compounds have proven to possess cytotoxic and antitumor activity in recent investigations and some have entered clinical trials and their effectiveness has been evaluated on humans.

These findings show the profound insight of Islamic physicians on cancer treatment. In spite of the lack of modern facilities and developed equipment, they introduced anticancer plants that have shown cytotoxic properties in new researches. The correlation between these findings signifies the originality of past experiences and studies, representing a worthwhile fund and valuable science dating back more than twelve centuries. This heritage is based on the experiences of thousands years of Greek, Indian and ancient Iranian physicians and relies on immense number of clinical trials on thousands of people. Furthermore, the application of traditional medicinal knowledge reinterpreted by modern data can lead to more effective and

evidence-based use of medicinal plants, which can contribute to therapeutic decisions on different illnesses.

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References

- [1] Jorjâni SI. *Zakhireh Khârazmshâhi (Treasure of Khârazmshâh)*. Photoprint of the manuscript dated 1206 AD, edited with Introduction by A.A. Saeedi Sirjani. Tehran: The Iranian Culture Foundation; 1977. pp. 358, 562-563, 585-586 (in Persian)
- [2] Razi MZ. *Al-Hâwi fi al-Tibbe (Continens)*. Toaymi H K (ed). Vol.4. Beirut: Dar Ihya al-Turath al-Arabi; 2002. pp. 5-16 (in Arabic)
- [3] Ibn Sina HA. *Al-Qanun fi al-Tibb (The Canon of Medicine)*. Vol.4. New Delhi: Jamia Hamdard; 1998. pp. 195-197(in Arabic)
- [4] Ahwazi Arjani AA. *Kamel al-Sina'ah al-Tibbiyah (The Perfect Art of the Medicine)* (in Arabic). Al-Dassuqi I (ed). Saadat Press; 1877. P.190-191 (in Arabic)
- [5] Qarshi I. *Al-Shamel fi al-Sinaat al-Tibbiyah (Comprehensive Book on the Art of Medicine)*. Ziedan Y (ed). Vol. 7. Abu Dhabi: al-Majma' al-Thaqafi (Publications of the Cultural Centre); 1999. pp. 943-944 (in Arabic)
- [6] Jorjâni SI. *Al-Aghrâz al-Tibbiyah wa al-Mabâhethi al-Alâiiyah (Medical goals and Alaâii's discussions)*. Photoprint of the manuscript dated 1399 AD, edited with Introduction by P.N. Khanlari. Tehran: The Iranian Culture Foundation; 1966. pp. 555-558 (in Persian)
- [7] Chashti MAK. *Exir-e-Azam*. 2 edn., Delhi: Nami Monshi Nolkshur; 1884. pp. 308-313 (in Persian)
- [8] Tonekaboni, M.M. *Tohfât al-Momenin (Rarity of the Faithful)*. Qum: Nour Vahy Press; 2011. pp.126-278, 328-423 (in Persian)

- [9] Aqili Khorasani MH. *Makhzan al-Adwiah (Drug Treasure)*. Reprinted from a copy which was printed in Calcutta dated in 1844. Tehran: Enqelab-e Eslami Publishing and Educational Organization; 1992. pp. 530-797 (in Persian)
- [10] Ibn Beytar Z. *Al-Jamee Le-Mofradaat al-Adwiah wal-Aghziyah (Comprehensive Book in Simple Drugs and Foods)*. Beirut: Dar- Al-Kotob Al-ilmiyah; 2001. pp. 160-185 (in Arabic)
- [11] al-Ghassani A. *Hadiqat Al-Azhar fi Mahiyat Al-'Ushb wa Al-'Uqar (Garden of Flowers in the Explanation of the Character of Herbs and Drugs)*. Beirut: Dar Al-Gharb Al-Islami; 1990. p.10 (in Arabic)
- [12] Razi MZ. *Taqasim Al-'Ilal (Kitab At-Taqsim Wa at-Tasgir)*. Hamami SM (ed). Aleppo: Aleppo University Publications, Institute for the History of Arabic Science; 1992. pp. 416-582 (in Arabic)
- [13] Ibn Beytar Z. *Al-Moghni fi-al-Tebb*. Digital photo of a handwritten manuscript dated 1834 AD. pp. 210-213 (in Arabic)
- [14] Tabari A. *Ferdows al-Hekmah fi al-Tibb (Paradise of Wisdom on Medicine)*. Al-Seddiqi (ed). Berlin: Aftab Press; 1928. pp.228-230 (in Arabic)
- [15] Akhawayni Bukhari AB. *Hedayat al-Mota'allemin fi al-Tibb (An Educational Guide for Medical Students)*. Matini J (ed). Mashhad: Ferdowsi University of Mashhad Publication; 1992. pp. 606-607 (in Arabic)
- [16] Lampronti I, Martello D, Bianchi N, Borgatti M, Lambertini E, Piva R, et al. In vitro antiproliferative effects on human tumor cell lines of extracts from the Bangladeshi medicinal plant *Aegle marmelos* Correa. *Phytomedicine*. 2003;10(4):300-8. PMID: 12809360
- [17] Costa-Lotufo LV, Khan MTH, Ather A, Wilke DV, Jimenez PC, Pessoa C, et al. Studies of the anticancer potential of plants used in Bangladeshi folk medicine. *J Ethnopharmacol*. 2005;99(1):21-30. PMID: 15848015
- [18] Khan MT, Lampronti I, Martello D, Bianchi N, Jabbar S, Choudhuri MS, et al. Identification of pyrogallol as an antiproliferative compound present in extracts from the medicinal plant *Emblica officinalis*: effects on in vitro cell growth of human tumor cell lines. *Int J Oncol*. 2002;21(1):187-92. PMID: 12063567
- [19] Hamid K, Diba F, Urmi KF, Uddin ME, Zohera FT, Habib MR. In vitro antioxidant and cytotoxicity screening of different bark extracts of *Aegle marmelos* L. *J Appl Pharmaceut Sci*. 2012;2(3):92-5.
- [20] Mustahil NA, Riyanto S, Sukari MA, Rahmani M, Mohd Nor SM, Ali AM. Antileukemic activity of extracts and constituents of *Aegle marmelos*. *Res J Chem Environ*. 2013;17(1):62-7.
- [21] Subramaniam D, Giridharan P, Murmu N, Shankaranarayanan NP, May R, Houchen CW, et al. Activation of apoptosis by 1-hydroxy-5,7-dimethoxy-2-naphthalene- car-

- boxaldehyde, a novel compound from *Aegle marmelos*. *Canc Res.* 2008;68(20):8573-81. doi: 10.1158/0008-5472.CAN-08-2372.
- [22] Abdullaev FI. Cancer chemopreventive and tumoricidal properties of saffron (*Crocus sativus* L.). *Exper Biol Med.* 2002;227(1):20-5. PMID: 11788779
- [23] Carotenuto A, Fattorusso E, Lanzotti V, Magno S, De Feo V, Carnuccio R, et al. Porri-
genins A and B, novel cytotoxic and antiproliferative sapogenins isolated from *Alli-
um porrum*. *J Natural Prod.* 1997;60(10):1003-7. PMID: 9358643
- [24] Fattorusso E, Lanzotti V, Tagliatela-Scafati O, Di Rosa M, Ianaro A. Cytotoxic sapo-
nins from bulbs of *Allium porrum* L. *J Agric Food Chem.* 2000;48(8):3455-62. PMID:
10956133
- [25] Lee KH, Kang HG, Cho CH, Lee MJ, Lee JH, Kim CH. Antimutagenic and antileuke-
mic activities of *Aloe vera* L. *Natural Prod Sci.* 2000;6(2):56-60.
- [26] Jose J, Sudhakaran S, Sumesh Kumar TM, Jayaraman S, Jayadevi Variyar E. A com-
parative evaluation of anticancer activities of flavonoids isolated from *Mimosa pudica*,
Aloe vera and *Phyllanthus niruri* against human breast carcinoma cell line (MCF-7) us-
ing MTT assay. *Int J Pharmacy Pharmaceut Sci.* 2014;6(2):319-22.
- [27] Lee KH, Kim JH, Lim DS, Kim CH. Anti-leukaemic and anti-mutagenic effects of
di(2-ethylhexyl)phthalate isolated from *Aloe vera* Linne. *J Pharmacy Pharmacol.*
2000;52(5):593-8. PMID: 10864149
- [28] Pecere T, Gazzola MV, Mucignat C, Parolin C, Vecchia FD, Cavaggioni A, et al. Aloe-
emodin is a new type of anticancer agent with selective activity against neuroecto-
dermal tumors. *Canc Res.* 2000;60(11):2800-4. PMID:10850417
- [29] Liu K, Park C, Li S, Lee KW, Liu H, He L, et al. Aloe-emodin suppresses prostate can-
cer by targeting the mTOR complex 2. *Carcinogenesis.* 2012 Jul;33(7):1406-11. PMID:
22532249. PMCID: 3405653.
- [30] Wasserman L, Avigad S, Beery E, Nordenberg J, Fenig E. The effect of aloe emodin
on the proliferation of a new Merkel carcinoma cell line. *Am J Dermatopathol.*
2002;24(1):17-22. PMID: 11803275
- [31] Acevedo-Duncan M, Russell C, Patel S, Patel R. Aloe-emodin modulates PKC iso-
zymes, inhibits proliferation, and induces apoptosis in U-373MG glioma cells. *Int Im-
munopharmacol.* 2004;4(14 SPEC.ISS.):1775-84. PMID:15531293
- [32] Ismail S, Haris K, Abdul Ghani ARI, Abdullah JM, Johan MF, Amohamed Yusoff
AA. Enhanced induction of cell cycle arrest and apoptosis via the mitochondrial
membrane potential disruption in human U87 malignant glioma cells by aloe emo-
din. *J Asian Natural Prod Res.* 2013;15(9):1003-12.

- [33] Lin JG, Chen GW, Li TM, Chouh ST, Tan TW, Chung JG. Aloe-Emodin induces apoptosis in T24 human bladder cancer cells through the p53 dependent apoptotic pathway. *J Urol*. 2006;175(1):343-7. PMID:16406939
- [34] Guo J, Xiao B, Liu Q, Gong Z, Le Y. Suppression of C-myc expression associates with anti-proliferation of aloe-emodin on gastric cancer cells. *Canc Invest*. 2008;26(4):369-74. doi: 10.1080/07357900701788130.
- [35] Hosseini J, Mahmoodi M, Jalili A, Hosseini S, Hosseini-Zijoud SM, Khoshdel A, et al. Aloe-emodin induces apoptosis through the up-regulation of fas in the human breast cancer cell line MCF-7. *Life Sci J*. 2014;11(SPEC.ISS.2):47-53.
- [36] Naveena, Bharath BK, Selvasubramanian. Antitumor activity of *Aloe vera* against Ehrlich ascitis carcinoma (EAC) in Swiss albino mice. *Int J Pharma Bio Sci*. 2011;2(2):400-9.
- [37] Grabarczyk H, Drozd B, Hladon B, Wojciechowska J. Sesquiterpene lactones. Part XV. New cytostatic active sesquiterpene lactone from herb of *Anthemis nobilis* L. *Polish J Pharmacol Pharmacy*. 1977;29(4):419-23. PMID: 905207
- [38] Kapadia GJ, Azuine MA, Rao GS, Arai T, Iida A, Tokuda H. Cytotoxic effect of the red beetroot (*Beta vulgaris* L.) extract compared to doxorubicin (adriamycin) in the human prostate (PC-3) and breast (MCF-7) cancer cell lines. *Anti-Canc Agents Medic Chem*. 2011;11(3):280-4. PMID: 21434853
- [39] Kapadia GJ, Rao GS, Ramachandran C, Iida A, Suzuki N, Tokuda H. Synergistic cytotoxicity of red beetroot (*Beta vulgaris* L.) extract with doxorubicin in human pancreatic, breast and prostate cancer cell lines. *J Complement Integ Med*. 2013;10. doi: 10.1515/jcim-2013-0007
- [40] Haroun AA, Taie HAA. Cytotoxicity and antioxidant activity of beta vulgaris extract released from grafted carbon nanotubes based nanocomposites. *Macromol Symp*. 2014;337(1):25-33. DOI: 10.1002/masy.201450303
- [41] Lee EJ, An D, Nguyen CTT, Patil BS, Kim J, Yoo KS. Betalain and betaine composition of greenhouse- or field-produced beetroot (*beta vulgaris* L.) and inhibition of HepG2 cell proliferation. *J Agric Food Chem*. 2014;62(6):1324-31. doi: 10.1021/jf404648u.
- [42] Zhao W, Entschladen F, Liu H, Niggemann B, Fang Q, Zaenker KS, et al. Boswellic acid acetate induces differentiation and apoptosis in highly metastatic melanoma and fibrosarcoma cells. *Canc Detect Prevention*. 2003;27(1):67-75. PMID: 12600419
- [43] Akihisa T, Tabata K, Banno N, Tokuda H, Nishihara R, Nakamura Y, et al. Cancer chemopreventive effects and cytotoxic activities of the triterpene acids from the resin of *Boswellia carteri*. *Biol Pharmaceut Bull*. 2006;29(9):1976-9. PMID: 16946522

- [44] Frank MB, Yang Q, Osban J, Azzarello JT, Saban MR, Saban R, et al. Frankincense oil derived from *Boswellia carteri* induces tumor cell specific cytotoxicity. *BMC Complement Alt Med*. 2009;9. doi: 10.1186/1472-6882-9-6.
- [45] Ali SI, Zhang CR, Mohamed AA, El-Baz FK, Hegazy AK, Kord MA, et al. Major constituents of *Boswellia carteri* resin exhibit cyclooxygenase enzyme inhibition and anti-proliferative activity. *Natural Prod Commun*. 2013;8(10):1365-6. PMID: 24354175
- [46] Bryant CS, Kumar S, Chamala S, Shah J, Pal J, Haider M, et al. Sulforaphane induces cell cycle arrest by protecting RB-E2F-1 complex in epithelial ovarian cancer cells. *Mol Canc*. 2010;9. doi: 10.1186/1476-4598-9-47
- [47] Melchini A, Traka MH. Biological profile of erucin: A new promising anticancer agent from cruciferous vegetables. *Toxins*. 2010;2(4):593-612. doi: 10.3390/toxins2040593.
- [48] Hafidh RR, Abdulmir AS, Abu Bakar F, Jalilian FA, Jahanshiri F, Abas F, et al. Novel anticancer activity and anticancer mechanisms of *Brassica oleracea* L. var. capitata f. rubra. *Eur J Integ Med*. 2013;5(5):450-64.
- [49] Thangam R, Suresh V, Rajkumar M, Vincent JD, Gunasekaran P, Anbazhagan C, et al. Antioxidant and in vitro anticancer effect of 2-pyrrolidinone rich fraction of *Brassica oleracea* var. capitata through induction of apoptosis in human cancer cells. *Phytother Res*. 2013;27(11):1664-70. doi: 10.1002/ptr.4908.
- [50] Kim SM, Park JH, Kim KD, Nam D, Shim BS, Kim SH, et al. Brassinin induces apoptosis in PC-3 human prostate cancer cells through the suppression of PI3K/Akt/mTOR/S6K1 signaling cascades. *Phytother Res*. 2014;28(3):423-31. doi: 10.1002/ptr.5010.
- [51] Ji YB, Dong F, Ma DB, Miao J, Jin LN, Liu ZF, et al. Optimizing the extraction of anti-tumor polysaccharides from the fruit of *Capparis spinosa* L. by response surface methodology. *Molecules*. 2012;17(6):7323-35. PMID: 22699566.
- [52] Yu L, Mo K, Wang W, Cui RT, Zou X, Ji YB. Relationship between apoptosis and [Ca²⁺]_i in HepG2 induced by *Capparis spinosa* alkaloid. *Chin Tradition Herbal Drugs*. 2009;40(2):255-8.
- [53] Loo WTY, Cheung MNB, Chow LWC. The inhibitory effect of a herbal formula comprising ginseng and *Carthamus tinctorius* on breast cancer. *Life Sci*. 2004;76(2):191-200. PMID: 15519364
- [54] Shi X, Ruan D, Wang Y, Ma L, Li M. Anti-tumor activity of safflower polysaccharide (SPS) and effect on cytotoxicity of CTL cells, NK cells of T739 lung cancer in mice. *Zhongguo Zhongyao Zazhi*. 2010;35(2):215-8. PMID: 20394298
- [55] Son ES, Lee JM, Choi JU, Park HR. Selective cytotoxicity of *Carthamus tinctorius* against glucose-deprived HT-29 human colon carcinoma cells. *J Medic Plant Res*. 2011;5(19):4931-7.

- [56] Sutton KM, Greenshields AL, Hoskin DW. Thymoquinone, a bioactive component of black caraway seeds, causes G1 phase cell cycle arrest and apoptosis in triple-negative breast cancer cells with mutant p53. *Nutri Canc*. 2014;66(3):408-18. doi: 10.1080/01635581.2013.878739.
- [57] Duraipandiyan V, Baskar AA, Ignacimuthu S, Muthukumar C, Al-Harbi NA. Anticancer activity of Rhein isolated from *Cassia fistula* L. flower. *Asian Pac J Trop Dis*. 2012;2(SUPPL.1):S517-S23. doi:10.1016/S2222-1808(12)60213-8
- [58] Linu M, Shankar S. In vitro cytotoxic activity of methanolic extract of stem bark of *Cassia fistula* L. *Res J Biotechnol*. 2012;7(4):141-8.
- [59] Gao XM, Shen YQ, Huang XZ, Yang LY, Shu LD, Hu QF, et al. 2"-ethyl-furanoflavone derivatives from the stems of *Cassia fistula* and their cytotoxicity. *J Brazil Chem Soc*. 2013;24(4):685-9.
- [60] Yang J, Wang H, Liu G, Lou J, Li L, Hu Q, et al. A new anthraquinone from the fruit of *Cassia fistula* and its cytotoxicity. *Asian J Chem*. 2014;26(14):4519-20.
- [61] Kumar S, Kapoor V, Gill K, Singh K, Xess I, Das SN, et al. Antifungal and antiproliferative protein from *Cicer arietinum*: A bioactive compound against emerging pathogens. *BioMed Res Int*. 2014;2014. doi: 10.1155/2014/387203. In Press.
- [62] Csupor-Löffler B, Hajdú Z, Réthy B, Zupkó I, Máthé I, Rédei T, et al. Antiproliferative activity of Hungarian asteraceae species against human cancer cell lines. Part II. *Phytother Res*. 2009;23(8):1109-15. doi: 10.1002/ptr.2755.
- [63] Zhou CX, Zou L, Zhao ZZ, Zhu H, He QJ, Yang B, et al. Terpenoids from *Cichorium intybus*. *Natural Prod Commun*. 2012;7(8):971-2. PMID: 22978206
- [64] Ka H, Park HJ, Jung HJ, Choi JW, Cho KS, Ha J, et al. Cinnamaldehyde induces apoptosis by ROS-mediated mitochondrial permeability transition in human promyelocytic leukemia HL-60 cells. *Canc Lett*. 2003;196(2):143-52. PMID: 12860272
- [65] Lee HS, Kim SY, Lee CH, Ahn YJ. Cytotoxic and mutagenic effects of *Cinnamomum cassia* bark-derived materials. *J Microbiol Biotechnol*. 2004;14(6):1176-81. doi: 10.1186/1471-2407-10-210.
- [66] Koppikar SJ, Choudhari AS, Suryavanshi SA, Kumari S, Chattopadhyay S, Kaul-Ghanekar R. Aqueous Cinnamon Extract (ACE-c) from the bark of *Cinnamomum cassia* causes apoptosis in human cervical cancer cell line (SiHa) through loss of mitochondrial membrane potential. *BMC Cancer*. 2010;10.
- [67] Kang HS, Ock J, Lee HJ, Lee YJ, Kwon BM, Hong SH. Early growth response protein 1 upregulation and nuclear translocation by 2'-benzoyloxycinnamaldehyde induces prostate cancer cell death. *Canc Lett*. 2013;329(2):217-27. doi: 10.1016/j.canlet.2012.11.006.

- [68] Ngoc TM, Nhiem NX, Khoi NM, Son DC, Hung TV, Kiem PV. A new coumarin and cytotoxic activities of constituents from *Cinnamomum cassia*. *Natural Prod Commun*. 2014;9(4):487-8. PMID: 24868863
- [69] Chinou I, Demetzos C, Harvala C, Roussakis C, Verbist JF. Cytotoxic and antibacterial labdane-type diterpenes from the aerial parts of *Cistus incanus* subsp. *creticus*. *Planta Medica*. 1994;60(1):34-6. PMID: 8134413
- [70] Dimas K, Demetzos C, Marsellos M, Sotiriadou R, Malamas M, Kokkinopoulos D. Cytotoxic activity of labdane type diterpenes against human leukemic cell lines in vitro. *Planta Medica*. 1998;64(3):208-11. PMID: 9581515
- [71] Dimas K, Demetzos C, Mitaku S, Vaos B, Marsellos M, Tzavaras T, et al. Cytotoxic activity and antiproliferative effects of a new semi-synthetic derivative of Ent-3 β -hydroxy-13-epi-manoyl oxide on human leukemic cell lines. *Anticanc Res*. 1999;19(5 B):4065-72. PMID: 11582537
- [72] Demetzos C, Dimas K, Hatziantoniou S, Anastasaki T, Angelopoulou D. Cytotoxic and anti-inflammatory activity of labdane and cis-clerodane type diterpenes. *Planta Medica*. 2001;67(7):614-8. PMID: 11582537
- [73] Skorić M, Todorović S, Gligorijević N, Janković R, Živković S, Ristić M, et al. Cytotoxic activity of ethanol extracts of in vitro grown *Cistus creticus* subsp. *creticus* L. on human cancer cell lines. *Indus Crops Prod*. 2012;38(1):153-9. doi:10.1016/j.indcrop.2012.01.017
- [74] Shah R, Gulati V, Palombo EA. Pharmacological properties of guggulsterones, the major active components of gum guggul. *Phytother Res*. 2012;26(11):1594-605. doi:10.1002/ptr.4647.
- [75] Jana S, Patra K, Sarkar S, Jana J, Mukherjee G, Bhattacharjee S, et al. Antitumorigenic potential of linalool is accompanied by modulation of oxidative stress: An in vivo study in sarcoma-180 solid tumor model. *Nutri Canc*. 2014;66(5):835-48. doi:10.1080/01635581.2014.904906.
- [76] Nithya TG, Sumalatha D. Evaluation of invitro anti-oxidant and anticancer activity of *Coriandrum Sativum* against human colon cancer HT- 29 cell lines. *Int J Pharmacy Pharmaceut Sci*. 2014;6(2):421-4.
- [77] Nair SC, Pannikar B, Panikkar KR. Antitumour activity of saffron (*Crocus sativus*). *Canc Lett*. 1991;57(2):109-14. PMID: 2025883
- [78] Tavakkol-Afshari J, Brook A, Mousavi SH. Study of cytotoxic and apoptogenic properties of saffron extract in human cancer cell lines. *Food Chem Toxicol*. 2008;46(11):3443-7. doi: 10.1016/j.fct.2008.08.018.
- [79] Samarghandian S, Boskabady MH, Davoodi S. Use of in vitro assays to assess the potential antiproliferative and cytotoxic effects of saffron (*Crocus sativus* L.) in human

- lung cancer cell line. *Pharmacognosy Mag.* 2010;6(24):309-14. doi: 10.4103/0973-1296.71799.
- [80] Escribano J, Alonso GL, Coca-Prados M, Fernández JA. Crocin, safranal and picrocrocin from saffron (*Crocus sativus* L.) inhibit the growth of human cancer cells in vitro. *Canc Lett.* 1996;100(1-2):23-30. PMID: 8620447
- [81] Noureini SK, Wink M. Antiproliferative effects of crocin in HepG2 cells by telomerase inhibition and hTERT down-regulation. *Asian Pac J Canc Prevention.* 2012;13(5):2305-9. PMID: 22901211
- [82] Malaekheh-Nikouei B, Mousavi SH, Shahsavand S, Mehri S, Nassirli H, Moallem SA. Assessment of cytotoxic properties of safranal and nanoliposomal safranal in various cancer cell lines. *Phytother Res.* 2013;27(12):1868-73. doi: 10.1002/ptr.4945.
- [83] Samarghandian S, Shabestari MM. DNA fragmentation and apoptosis induced by safranal in human prostate cancer cell line. *Ind J Urol.* 2013;29(3):177-83. doi: 10.4103/0970-1591.117278.
- [84] Geromichalos GD, Papadopoulos T, Sahpazidou D, Sinakos Z. Safranal, a *Crocus sativus* L constituent suppresses the growth of K-562 cells of chronic myelogenous leukemia. In silico and in vitro study. *Food Chem Toxicol.* 2014;74:45-50. doi: 10.1016/j.fct.2014.09.001.
- [85] Kim SH, Lee JM, Kim SC, Park CB, Lee PC. Proposed cytotoxic mechanisms of the saffron carotenoids crocin and crocetin on cancer cell lines. *Biochem Cell Biol.* 2014;92(2):105-11. doi: 10.1139/bcb-2013-0091.
- [86] Rezaee R, Jamialahmadi K, Zanjani BR, Mahmoudi M, Abnous K, Rabe SZT, et al. Crocin effects on human myeloma cells regarding intracellular redox state, DNA fragmentation, and apoptosis or necrosis profile. *Jundishapur J Natural Pharmaceut Prod.* 2014;9(4). PMID: 25625054
- [87] Rastgoo M, Hosseinzadeh H, Alavizadeh H, Abbasi A, Ayati Z, Jaafari MR. Antitumor activity of PEGylated nanoliposomes containing crocin in mice bearing C26 colon carcinoma. *Planta Medica.* 2013;79(6):447-51. doi: 10.1055/s-0032-1328363.
- [88] Begum Y, Hossen F, Rahman MM. In vitro cytotoxic properties of methanol extracts of the seeds of *benincasa hispida* and *cucurbita maxima*. *Eur J Sci Res.* 2011;54(2):193-7.
- [89] Saha P, Mazumder UK, Haldar PK, Naskar S, Kundu S, Bala A, et al. Anticancer activity of methanol extract of *Cucurbita maxima* against Ehrlich as-cites carcinoma. *Int J Res Pharmaceut Sci.* 2011;2(1):52-9.
- [90] Kikuchi T, Takebayashi M, Shinto M, Yamada T, Tanaka R. Three new multiflorane-type triterpenes from pumpkin (*Cucurbita maxima*) seeds. *Molecules.* 2013;18(5):5568-79. doi: 10.3390/molecules18055568.

- [91] Shokrzadeh M, Azadbakht M, Ahangar N, Hashemi A, Saeedi Saravi SS. Cytotoxicity of hydro-alcoholic extracts of *Cucurbita pepo* and *Solanum nigrum* on HepG2 and CT26 cancer cell lines. *Pharmacognosy Mag.* 2010;6(23):176-9. doi: 10.4103/0973-1296.66931
- [92] Jafarian A, Ghannadi A, Mohebi B. Cytotoxic effects of chloroform and hydroalcoholic extracts of aerial parts of *Cuscuta chinensis* and *Cuscuta epithimum* on Hela, HT29 and MDA-MB-468 tumor cells. *Res Pharmaceut Sci.* 2014;9(2):115-22.
- [93] Attard E, Cuschieri A. Cytotoxicity of Cucurbitacin E extracted from *Ecballium elaterium* in vitro. *J Natural Remedies.* 2004;4(2):137-44.
- [94] Attard E, Cuschieri A, Brincat MP. Morphological effects induced by Cucurbitacin E on ovarian cancer cells in vitro. *J Natural Remedies.* 2005;5(1):70-4.
- [95] Bohlooli S, Jafari N, Jahed S. Cytotoxic effect of freeze-dried extract of *Ecballium elaterium* fruit on gastric adenocarcinoma (AGS) and esophageal squamous cell carcinoma (KYSE30) cell lines. *J Gastrointest Canc.* 2012;43(4):579-83. doi: 10.1007/s12029-012-9383-4.
- [96] Jacquot C, Rousseau B, Carbonnelle D, Chinou I, Malleter M, Tomasoni C, et al. Cucurbitacin-D-induced CDK1 mRNA up-regulation causes proliferation arrest of a non-small cell lung carcinoma cell line (NSCLC-N6). *Anticanc Res.* 2014;34(9):4797-806. PMID: 25202060
- [97] Khodarahmi GA, Ghasemi N, Hassanzadeh F, Safaie M. Cytotoxic effects of different extracts and latex of *ficus carica* L. on HeLa cell line. *Iranian J Pharmaceut Res.* 2011;10(2):273-7. PMID: 24250354
- [98] Hamta A, Shariatzadeh SMA, Soleimani Mehranjani M, Fallah Huseini H, Hosseina-badi F. The cytotoxic effects of *Glycyrrhiza glabra* L. root extract on 4T1 cell line derived from BALB/c mice mammary tumors. *J Medic Plants.* 2014;13(50):92-103.
- [99] Dorn DC, Alexenizer M, Hengstler JG, Dorn A. Tumor cell specific toxicity of *Inula helenium* extracts. *Phytother Res.* 2006;20(11):970-80. PMID: 16912983
- [100] Mazzio E, Badisa R, Mack N, Deiab S, Soliman KFA. High throughput screening of natural products for anti-mitotic effects in MDA-MB-231 human breast carcinoma cells. *Phytother Res.* 2014;28(6):856-67. doi: 10.1002/ptr.5065.
- [101] Rahman MA, Hasan MS, Hossain MA, Biswas NN. Analgesic and cytotoxic activities of *Jasminum sambac* (L.) Aiton. *Pharmacologyonline.* 2011;1:124-31.
- [102] Liu JX, Di DL, Wei XN, Han Y. Cytotoxic diarylheptanoids from the pericarps of walnuts (*Juglans regia*). *Planta Medica.* 2008;74(7):754-9. doi: 10.1055/s-2008-1074536.
- [103] Salimi M, Majd A, Sepahdar Z, Azadmanesh K, Irian S, Ardestaniyan MH, et al. Cytotoxicity effects of various *Juglans regia* (walnut) leaf extracts in human cancer cell lines. *Pharmaceut Biol.* 2012;50(11):1416-22. doi: 10.3109/13880209.2012.682118.

- [104] Li CY, Du HJ, Su XH, Zhong YJ, Yuan ZP, Li YF, et al. Juglanones A and B: Two novel tetralone dimers from walnut pericarp (*Juglans regia*). *Helvetica Chim Acta*. 2013;96(6):1031-5. DOI: 10.1002/hlca.201200525
- [105] Salimi M, Ardestaniyan MH, Mostafapour Kandelous H, Saeidnia S, Gohari AR, Amanzadeh A, et al. Anti-proliferative and apoptotic activities of constituents of chloroform extract of *Juglans regia* leaves. *Cell Prolifer*. 2014;47(2):172-9.
- [106] Han YF, Cao GX, Gao XJ, Xia M. Isolation and characterisation of the sesquiterpene lactones from *Lactuca sativa* L var. *anagustata*. *Food Chemistry*. 2010;120(4):1083-8. doi:10.1016/j.foodchem.2009.11.056
- [107] Kivçak B, Mert T. Preliminary evaluation of cytotoxic properties of *Laurus nobilis* leaf extracts. *Fitoterapia*. 2002;73(3):242-3. PMID: 12048018
- [108] Dall'Acqua S, Viola G, Giorgetti M, Loi MC, Innocenti G. Two new sesquiterpene lactones from the leaves of *Laurus nobilis*. *Chem Pharmaceut Bull*. 2006;54(8):1187-9. PMID:16880666
- [109] Barla A, Topçu G, Öksüz S, Tümen G, Kingston DGI. Identification of cytotoxic sesquiterpenes from *Laurus nobilis* L. *Food Chem*. 2007;104(4):1478-84. doi:10.1016/j.foodchem.2007.02.019
- [110] Loizzo MR, Tundis R, Menichini F, Saab AM, Statti GA. Cytotoxic activity of essential oils from Labiatae and Lauraceae families against in vitro human tumor models. *Anticanc Res*. 2007;27(5 A):3293-9. PMID: 17970073
- [111] Li Q, Gao W, Cao J, Bi X, Chen G, Zhang X, et al. New cytotoxic compounds from flowers of *Lawsonia inermis* L. *Fitoterapia*. 2014;94:148-54.
- [112] Lehraiki A, Attoumbré J, Bienaimé C, Matifat F, Bensaddek L, Nava-Saucedo E, et al. Extraction of lignans from flaxseed and evaluation of their biological effects on breast cancer MCF-7 and MDA-MB-231 cell lines. *J Medic Food*. 2010;13(4):834-41. doi: 10.1089/jmf.2009.0172.
- [113] de Sousa AC, Alviano DS, Blank AF, Barreto Alves P, Alviano CS, Gattass CR. *Melissa officinalis* L. essential oil: Antitumoral and antioxidant activities. *J Pharmacy Pharmacol*. 2004;56(5):677-81. PMID:15142347
- [114] Encalada MA, Hoyos KM, Rehecho S, Berasategi I, de Ciriano MGI, Ansorena D, et al. Anti-proliferative Effect of *Melissa officinalis* on Human Colon Cancer Cell Line. *Plant Foods Human Nutri*. 2011;66(4):328-34. doi: 10.1007/s11130-011-0256-y.
- [115] Saraydin SU, Tuncer E, Tepe B, Karadayi S, Özer H, Şen M, et al. Antitumoral effects of *Melissa officinalis* on breast cancer in vitro and in vivo. *Asian Pac J Canc Prevention*. 2012;13(6):2765-70. PMID: 22938456

- [116] De Queiroz RM, Takiya CM, Guimarães LPTP, Rocha GDG, Alviano DS, Blank AF, et al. Apoptosis-inducing effects of *Melissa officinalis* L. essential oil in glioblastoma multiforme cells. *Canc Invest*. 2014;32(6):226-35. doi: 10.3109/07357907.2014.905587
- [117] Cottiglia F, Casu L, Leonti M, Caboni P, Floris C, Busonera B, et al. Cytotoxic phloroglucinols from the leaves of *Myrtus communis*. *J Natural Prod*. 2012;75(2):225-9. doi: 10.1021/np2009219.
- [118] Youssef DTA, Khalifa AA. Cytotoxic quaternary alkaloids from the flowers of *Narcissus tazetta*. *Pharmazie*. 2001;56(10):818-22. PMID: 11683132
- [119] Liu J, Li Y, Ren W, Hu WX. Apoptosis of HL-60 cells induced by extracts from *Narcissus tazetta* var. *chinensis*. *Canc Lett*. 2006;242(1):133-40. PMID: 16427186
- [120] Juan ME, Wenzel U, Daniel H, Planas JM. Erythrodiol, a natural triterpenoid from olives, has antiproliferative and apoptotic activity in HT-29 human adenocarcinoma cells. *Mol Nutri Food Res*. 2008;52(5):595-9. doi: 10.1002/mnfr.200700300.
- [121] Bouallagui Z, Han J, Isoda H, Sayadi S. Hydroxytyrosol rich extract from olive leaves modulates cell cycle progression in MCF-7 human breast cancer cells. *Food Chem Toxicol*. 2011;49(1):179-84. doi: 10.1016/j.fct.2010.10.014.
- [122] Reyes-Zurita FJ, Pachón-Peña G, Lizárraga D, Rufino-Palomares EE, Cascante M, Lupiáñez JA. The natural triterpene maslinic acid induces apoptosis in HT29 colon cancer cells by a JNK-p53-dependent mechanism. *BMC Cancer*. 2011;11. doi: 10.1186/1471-2407-11-154.
- [123] Busnena BA, Foudah AI, Melancon T, El Sayed KA. Olive secoiridoids and semisynthetic bioisostere analogues for the control of metastatic breast cancer. *Bioorg Medic Chem*. 2013;21(7):2117-27. doi: 10.1016/j.bmc.2012.12.050.
- [124] University of Southern California. *Study of Noscapine for Patients With Low Grade Non Hodgkin's Lymphoma or Chronic Lymphocytic Leukemia Refractory to Chemotherapy*. In: ClinicalTrials.gov [Internet]. Bethesda (MD): National Library of Medicine (US). 2015 Jan 25. Available from: <http://clinicaltrials.gov/ct2/results?term=noscapine&Search=Search>.
- [125] Zhang CH, Wang ZT, Yang YP, Sun QS. A novel cytotoxic neophysalin from *Physalis alkekengi* var. *francheti*. *Chin Chem Lett*. 2009;20(11):1327-30.
- [126] He H, Zang LH, Feng YS, Chen LX, Kang N, Tashiro SI, et al. Physalin A induces apoptosis via p53-Noxa-mediated ROS generation, and autophagy plays a protective role against apoptosis through p38-NF- κ B survival pathway in A375-S2 cells. *J Ethnopharmacol*. 2013;148(2):544-55. doi: 10.1016/j.jep.2013.04.051.
- [127] He H, Zang LH, Feng YS, Wang J, Liu WW, Chen LX, et al. Physalin a induces apoptotic cell death and protective autophagy in HT1080 human fibrosarcoma cells. *J Natural Prod*. 2013;76(5):880-8. doi: 10.1021/np400017k.

- [128] Rezaei PF, Fouladdel S, Hassani S, Yousefbeyk F, Ghaffari SM, Amin G, et al. Induction of apoptosis and cell cycle arrest by pericarp polyphenol-rich extract of Baneh in human colon carcinoma HT29 cells. *Food Chem Toxicol.* 2012;50(3-4):1054-9. doi: 10.1016/j.fct.2011.11.012.
- [129] Balan KV, Demetzos C, Prince J, Dimas K, Cladaras M, Han Z, et al. Induction of apoptosis in human colon cancer HCT116 cells treated with an extract of the plant product, chios mastic gum. *In Vivo.* 2005;19(1):93-102. PMID: 15796160
- [130] Chiang LC, Chiang W, Chang MY, Lin CC. In vitro cytotoxic, antiviral and immunomodulatory effects of *Plantago major* and *Plantago asiatica*. *Am J Chin Med.* 2003;31(2): 225-34. PMID: 12856861
- [131] Gálvez M, Martín-Cordero C, López-Lázaro M, Cortés F, Ayuso MJ. Cytotoxic effect of *Plantago spp.* on cancer cell lines. *J Ethnopharmacol.* 2003;88(2-3):125-30. PMID: 12963131
- [132] Mitrocotsa D, Bosch S, Mitaku S, Dimas C, Skaltsounis AL, Harvala C, et al. Cytotoxicity against human leukemic cell lines, and the activity on the expression of resistance genes of flavonoids from *Platanus orientalis*. *Anticanc Res.* 1999;19(3 A):2085-8. PMID:10470152
- [133] Mitrokotsa D, Mitaku S, Demetzos C, Harvala C, Mentis A, Perez S, et al. Bioactive compounds from the buds of *Platanus orientalis* and isolation of a new kaempferol glycoside. *Planta Medica.* 1993;59(6):517-20. PMID: 8302950
- [134] Dimas K, Demetzos C, Mitaku S, Marselos M, Tzavaras T, Kokkinopoulos D. Cytotoxic activity of kaempferol glycosides against human leukaemic cell lines in vitro. *Pharmacol Res.* 2000;41(1):85-8. PMID: 10600274
- [135] Afifi-Yazar FU, Kasabri V, Abu-Dahab R. Medicinal plants from Jordan in the treatment of cancer: Traditional uses vs in vitro and in vivo evaluations part 1. *Planta Medica.* 2011;77(11):1203-9. doi: 10.1055/s-0030-1270832.
- [136] Khan I, Sangwan PL, Dar AA, Rafiq RA, Farrukh MR, Dhar JK, et al. A validated high-performance thin-layer chromatography method for the identification and simultaneous quantification of six markers from *Platanus orientalis* and their cytotoxic profiles against skin cancer cell lines. *J Separ Sci.* 2013;36(16):2602-10. doi: 10.1002/jssc.201300380.
- [137] Habibi Roudkenar M, Mohammadi Roushandeh A, Delazar A, Halabian R, Soleimani Rad J, Mehdipour A, et al. Effects of polygonum aviculare herbal extract on proliferation and apoptotic gene expression of MCF-7. *DARU, J Pharmaceut Sci.* 2011;19(5): 326-31. PMID:
- [138] Xin HL, Xu YF, Hou YH, Zhang YN, Yue XQ, Lu JC, et al. Two novel triterpenoids from *Portulaca oleracea* L. *Helvetica Chim Acta.* 2008;91(11):2075-80.
- [139] Tan GCS, Wong KM, Pearle-Wong GQ, Yeo SL, Yeap SK, Yiap BC, et al. In vitro cytotoxic and antiproliferative effects of *portulaca oleracea* methanol extract on breast, cer-

- vical, colon and nasopharyngeal cancerous cell lines. *Sains Malaysiana*. 2013;42(7):927-35.
- [140] Farshori NN, Al-Sheddi ES, Al-Oqail MM, Musarrat J, Al-Khedhairi AA, Siddiqui MA. Cytotoxicity assessments of *Portulaca oleracea* and *Petroselinum sativum* seed extracts on human hepatocellular carcinoma cells (HepG2). *Asian Pac J Canc Prevention*. 2014;15(16):6633-8. PMID: 25169500
- [141] Tian JL, Liang X, Gao PY, Li DQ, Sun Q, Li LZ, et al. Two new alkaloids from *Portulaca oleracea* and their cytotoxic activities. *J Asian Natural Prod Res*. 2014;16(3):259-64. doi: 10.1080/10286020.2013.866948.
- [142] Jeune MAL, Kumi-Diaka J, Brown J. Anticancer activities of pomegranate extracts and genistein in human breast cancer cells. *J Medic Food*. 2005;8(4):469-75. PMID: 16379557
- [143] Banerjee N, Talcott S, Safe S, Mertens-Talcott SU. Cytotoxicity of pomegranate polyphenolics in breast cancer cells in vitro and vivo: Potential role of miRNA-27a and miRNA-155 in cell survival and inflammation. *Breast Canc Res Treat*. 2012;136(1):21-34. doi: 10.1007/s10549-012-2224-0.
- [144] Jayakumar S, Haridass S, Krishnamurthy V. Anticancer activity of *Punica Granatum* rind extracts against human lung cancer cell line. *Asian J Pharmaceut Clin Res*. 2012;5(SUPPL 2):204-10.
- [145] Joseph MM, Aravind SR, Varghese S, Mini S, Sreelekha TT. Evaluation of antioxidant, antitumor and immunomodulatory properties of polysaccharide isolated from fruit rind of *Punica granatum*. *Mol Med Rep*. 2012;5(2):489-96. doi: 10.3892/mmr.2011.638.
- [146] Bekir J, Mars M, Vicendo P, Fterrich A, Bouajila J. Chemical composition and antioxidant, anti-inflammatory, and antiproliferation activities of pomegranate (*Punica granatum*) flowers. *J Medic Food*. 2013;16(6):544-50. doi: 10.1089/jmf.2012.0275.
- [147] Joseph MM, Aravind SR, George SK, Varghese S, Sreelekha TT. A galactomannan polysaccharide from *Punica granatum* imparts in vitro and in vivo anticancer activity. *Carbohydr Polymers*. 2013;98(2):1466-75. doi: 10.1016/j.carbpol.2013.07.023.
- [148] Salah-Abbes JB, Abbès S, Abdel-Wahhab MA, Oueslati R. In-vitro free radical scavenging, antiproliferative and anti-zearalenone cytotoxic effects of 4-(methylthio)-3-butenyl isothiocyanate from Tunisian *Raphanus sativus*. *J Pharmacy Pharmacol*. 2010;62(2):231-9. PMID: 20487203
- [149] Kim KH, Moon E, Kim SY, Choi SU, Lee JH, Lee KR. 4-Methylthio-butanyl derivatives from the seeds of *Raphanus sativus* and their biological evaluation on anti-inflammatory and antitumor activities. *J Ethnopharmacol*. 2014;151(1):503-8. doi: 10.1016/j.jep.2013.11.003.

- [150] Zarai Z, Chobba IB, Mansour RB, Békir A, Gharsallah N, Kadri A. Essential oil of the leaves of *Ricinus communis* L.: In vitro cytotoxicity and antimicrobial properties. *Lipids Health Dis.* 2012;11. doi: 10.1186/1476-511X-11-102.
- [151] Rezaie-Tavirani M, Fayazfar S, Heydari-Keshel S, Rezaee MB, Zamanian-Azodi M, Rezaei-Tavirani M, et al. Effect of essential oil of *Rosa damascena* on human colon cancer cell line SW742. *Gastroenterol Hepatol Bed to Bench.* 2013;6(1):25-31. PMID: 24834241
- [152] Kuo YH, Hsu YW, Liaw CC, Lee JK, Huang HC, Kuo LMY. Cytotoxic phenylpropanoids glycosides from the stems of *Smilax china*. *J Natural Prod.* 2005;68(10):1475-8. PMID: 16252910
- [153] Li YL, Gan GP, Zhang HZ, Wu HZ, Li CL, Huang YP, et al. A flavonoid glycoside isolated from *Smilax china* L. rhizome in vitro anticancer effects on human cancer cell lines. *J Ethnopharmacol.* 2007;113(1):115-24. PMID: 17606345
- [154] Wu LS, Wang XJ, Wang H, Yang HW, Jia AQ, Ding Q. Cytotoxic polyphenols against breast tumor cell in *Smilax china* L. *J Ethnopharmacol.* 2010;130(3):460-4. PMID: 20669365
- [155] Hu K, Kobayashi H, Dong A, Jing Y, Iwasaki S, Yao X. Antineoplastic agents III: Steroidal glycosides from *solanum nigrum*. *Planta Medica.* 1999;65(1):35-8. PMID: 10083842
- [156] Zhou X, He X, Wang G, Gao H, Zhou G, Ye W, et al. Steroidal saponins from *Solanum nigrum*. *J NaturProd.* 2006;69(8):1158-63. PMID: 16933867:
- [157] Heo KS, Lee SJ, Ko JH, Lim K, Lim KT. Glycoprotein isolated from *Solanum nigrum* L. inhibits the DNA-binding activities of NF- κ B and AP-1, and increases the production of nitric oxide in TPA-stimulated MCF-7 cells. *Toxicol In Vitro.* 2004;18(6):755-63. PMID: 15465640
- [158] Heo KS, Lee SJ, Lim KT. Cytotoxic effect of glycoprotein isolated from *Solanum nigrum* L. through the inhibition of hydroxyl radical-induced DNA-binding activities of NF-kappa B in HT-29 cells. *Environ Toxicol Pharmacol.* 2004;17(1):45-54. doi: 10.1016/j.etap.2004.02.003.
- [159] Lee SJ, Oh PS, Ko JH, Lim K, Lim KT. A 150-kDa glycoprotein isolated from *Solanum nigrum* L. has cytotoxic and apoptotic effects by inhibiting the effects of protein kinase C alpha, nuclear factor-kappa B and inducible nitric oxide in HCT-116 cells. *Canc Chemother Pharmacol.* 2004;54(6):562-72. PMID: 15349752
- [160] Lim KT. Glycoprotein isolated from *Solanum nigrum* L. kills HT-29 cells through apoptosis. *J Medic Food.* 2005;8(2):215-26. PMID: 16117614
- [161] Patel S, Gheewala N, Suthar A, Shah A. In-vitro cytotoxicity activity of *Solanum nigrum* extract against Hela cell line and Vero cell line. *Int J Pharmacy Pharmaceut Sci.* 2009;1(SUPPL. 1):38-46.

- [162] Huang HC, Syu KY, Lin JK. Chemical composition of *Solanum nigrum* linn extract and induction of autophagy by leaf water extract and its major flavonoids in AU565 breast cancer cells. *J Agric Food Chem*. 2010;58(15):8699-708. doi: 10.1021/jf101003v.
- [163] Sun L, Zhao Y, Li X, Yuan H, Cheng A, Lou H. A lysosomal-mitochondrial death pathway is induced by solamargine in human K562 leukemia cells. *Toxicol In Vitro*. 2010;24(6):1504-11. doi: 10.1016/j.tiv.2010.07.013. Epub 2010 Jul 18.
- [164] Akbar N, Thakur VS, Yunus M, Mahdi AA, Gupta S. Selective cell cycle arrest and induction of apoptosis in human prostate cancer cells by a polyphenol-rich extract of *Solanum nigrum*. *Int J Mol Med*. 2012;29(2):277-84. PMID: 22076244
- [165] Gabrani R, Jain R, Sharma A, Sarethy IP, Dang S, Gupta S. Antiproliferative effect of *Solanum nigrum* on human leukemic cell lines. *Ind J Pharmaceut Sci*. 2012;74(5):451-3. PMID: 23716874
- [166] Mathema VB, Koh YS, Thakuri BC, Sillanpää M. Parthenolide, a sesquiterpene lactone, expresses multiple anti-cancer and anti-inflammatory activities. *Inflammation*. 2012;35(2):560-5. PMID: 21603970
- [167] Lee SH, Ryu SY, Choi SU, Lee CO, No Z, Kim SK, et al. Hydrolysable tannins and related compound having cytotoxic activity from the fruits of *Terminalia chebula*. *Arch Pharmacol Res*. 1995;18(2):118-20.
- [168] Saleem A, Husheem M, Härkönen P, Pihlaja K. Inhibition of cancer cell growth by crude extract and the phenolics of *Terminalia chebula* retz. fruit. *J Ethnopharmacol*. 2002;81(3):327-36. PMID: 12127233
- [169] Alsemari A, Alkhodairy F, Aldakan A, Al-Mohanna M, Bahoush E, Shinwari Z, et al. The selective cytotoxic anti-cancer properties and proteomic analysis of *Trigonella Foenum-Graecum*. *BMC Complement Alt Med*. 2014;14. PMID: 24679057
- [170] Gerlach SL, Rathinakumar R, Chakravarty G, Göransson U, Wimley WC, Darwin SP, et al. Anticancer and chemosensitizing abilities of cycloviolacin 02 from *Viola odorata* and psyle cyclotides from *Psychotria leptothyrsa*. *Biopolymers*. 2010;94(5):617-25. PMID: 20564026
- [171] Berardi V, Ricci F, Castelli M, Galati G, Risuleo G. Resveratrol exhibits a strong cytotoxic activity in cultured cells and has an antiviral action against polyomavirus: Potential clinical use. *J Exper Clin Canc Res*. 2009;28(1). PMID:19570215
- [172] Aghbali A, Hosseini SV, Delazar A, Gharavi NK, Shahneh FZ, Orangi M, et al. Induction of apoptosis by grape seed extract (*Vitis vinifera*) in oral squamous cell carcinoma. *Bosnian J Basic Med Sci*. 2013;13(3):186-91. PMID: 23988171
- [173] Giovannelli L, Innocenti M, Santamaria AR, Bigagli E, Pasqua G, Mulinacci N. Antitumoural activity of viniferin-enriched extracts from *Vitis vinifera* L. cell cultures. *Natur Prod Res*. 2014;28(22):2006-16. PMID: 24949881

- [174] Hamadani SF, Pourseyedi S, Esmaeili-Mahani S. Cytotoxic effect of Rishbaba grape (*Vitis vinifera*) seed extract on human breast cancer cell line (MCF-7) and its interaction with the anticancer drug doxorubicin. *Physiol Pharmacol*. 2014;18(2):170-8.
- [175] Keum YS, Kim J, Lee KH, Park KK, Surh YJ, Lee JM, et al. Induction of apoptosis and caspase-3 activation by chemopreventive [6]-paradol and structurally related compounds in KB cells. *Canc Lett*. 2002;177(1):41-7. PMID: 11809529
- [176] Wang CC, Chen LG, Lee LT, Yang LL. Effects of 6-Gingerol, an antioxidant from ginger, on inducing apoptosis in human leukemic HL-60 cells. *In Vivo*. 2003;17(6):641-5. PMID:14758732
- [177] Wei QY, Ma JP, Cai YJ, Yang L, Liu ZL. Cytotoxic and apoptotic activities of diarylheptanoids and gingerol-related compounds from the rhizome of Chinese ginger. *J Ethnopharmacol*. 2005;102(2):177-84. PMID: 16024193
- [178] Nigam N, Bhui K, Prasad S, George J, Shukla Y. [6]-Gingerol induces reactive oxygen species regulated mitochondrial cell death pathway in human epidermoid carcinoma A431 cells. *Chemico-Biol Interact*. 2009;181(1):77-84. PMID: 19481070
- [179] Yang L, Zhou C, Huang K, Song L, Zheng Q, Yu R, et al. Antioxidative and cytotoxic properties of diarylheptanoids isolated from *Zingiber officinale*. *Zhongguo Zhongyao Zazhi*. 2009;34(3):319-23. PMID: 19445158
- [180] Liu Q, Peng YB, Qi LW, Cheng XL, Xu XJ, Liu LL, et al. The cytotoxicity mechanism of 6-shogaol-treated HeLa human cervical cancer cells revealed by label-free shotgun proteomics and bioinformatics analysis. *Evidence-based Complementary and Alternative Medicine*. 2012;2012. PMID: 23243437
- [181] Lv L, Chen H, Soroka D, Chen X, Leung T, Sang S. 6-gingerdiols as the major metabolites of 6-gingerol in cancer cells and in mice and their cytotoxic effects on human cancer cells. *J Agric Food Chem*. 2012;60(45):11372-7. PMID: 23066935