

Plant Archives

Journal homepage: http://www.plantarchives.org doi link : https://doi.org/10.51470/PLANTARCHIVES.2021.v21.S1.398

REVIEW ARTICLE: GARLIC: NATURE'S PROTECTION AGAINST WOUNDS

Gurjot Kaur and Reena Gupta*

Lovely institute of technology, Pharmacology department, Lovely professional university, Jalandhar-Delhi G.T. Road(NH-1), Phagwara, Punjab (India)-144411

*Corresponding Author E-mail: reena.ph14@gmail.com

ABSTRACT

Wounds can occur due to any injury, disease state or insect bite or abrasions. Wound healing is a complex process that includes many phases and involves a number of inflammatory mediators wound can be classified as acute and chronic. Other than this wound can also be classified on the basis of layer of skin that gets damaged: Superficial wound, Partial thickness wound and Full thickness wound. Garlic (*Allium sativum*) has been used as food and also used for the treatment of many diseases, for healing of wounds etc. Garlic is perennial liliaceous herb with a bulb having organosulphur compounds like Diallyl disulphide, diallyl sulphide, diallyl trisulphide and sulphur dioxide others are allin, allicin, etc. Garlic having major properties includes anti-cancer, against cardiovascular disease, anti-oxidant,anti-microbial, in wound healing etc.

Introduction

Skin act as protective barrier for the body which protects the body from external environment. Epidermal layer of skin can be damaged due to injury that subsequently leads to the formation of the wound. Wounds can occur due to any injury, disease state or insect bite or abrasions. These are defined as the breakage in the regular continuality or integrity of tissue. Subsequent to the wound injury there is inflammation followed by infection in the wounded area. Invasion by various pathogenic microorganisms at the wounded tissue results in severe infection of the wounded surface. Anatomical and functional integrity of tissue get disrupted due to chronic wound infection. The humoral and cellular components of body play an active role in the pathogenesis of wound infection (Steven Percival, Keith Cutting).

The most commonly seen wounds in humans are the lower extremity wounds in diabetic patients, wounds following extensive necrosis followed by infection and the chronic wounds (Hermans, M.H.E. and T. Treadwell)

Wound healing is a complex process that includes many phases and involves a number of inflammatory mediators. Complex biochemical events such as inflammation, coagulation, homeostasis, proliferation and granulation tissue formation are involved in wound healing. These events are divided into three phases: inflammatory, proliferative and remodelling phase. But different endogenous and exogenous factors such as microbial infection on wound, poor blood supply to the wound area and nutritional deficiency such as vitamin C and Zinc results in delay in healing of wound (Thomas *et al.*, 2011). On the basis of mechanism and duration of healing, wound can be classified as acute and chronic. Other than this wound can also be classified on the basis of layer of skin that gets damaged: Superficial wound, Partial thickness wound and Full thickness wound. Currently a wide range of topical antibiotics, antifungal and antiseptic drugs are used for the treatment of wound infections. Other than these new effective techniques such as tissue engineered growth technique, recombinant growth factors and silver dressings are also used nowadays for the treatment of wound infections. A number of cleansing solutions and gels are also used for rinsing and cleaning of wound such as sterile normal saline, sterile water, Povidone iodine, Poly-hexane biguanide and sodium hypochlorite (Benjamin A. Lipsky1,2 and Christopher Hoey).

Marketed antimicrobial drugs used for the treatment of wound infections are associated with various side effects like itching, redness of skin and blister formation. Resistant cases with the use of topical antibiotics and antifungal are also rising nowadays. This led to failure of antimicrobial therapy used to treat wound infection. Other than this major drawback associated with new advanced effective techniques such as recombinant growth factor, silver dressings is that these are very expensive (Tiwari and Jain, 2011).

Considering the limitations associated with the commercially available antibiotics and newer techniques, there is urgent need to develop as safer and cost-effective wound healing agent. Medicinal plants have been used from the ancient time for the treatment of various ailments. Large money has been invested by the pharmaceutical companies to isolate the essential Phyto-constituents from the medicinal plants to use them as efficacious medicinal and

cosmeceuticals agents. A number of medicinal plants which contain alkaloids and flavonoids as chief chemical constituents can effectively enhance the wound healing process. Medicinal plants such as Neem, Aloe Vera, Tulsi are rich source of alkaloids and flavonoids, so such plants can strongly enhance the wound healing process.

The present review article summarizes the role of garlic on the wound healing process

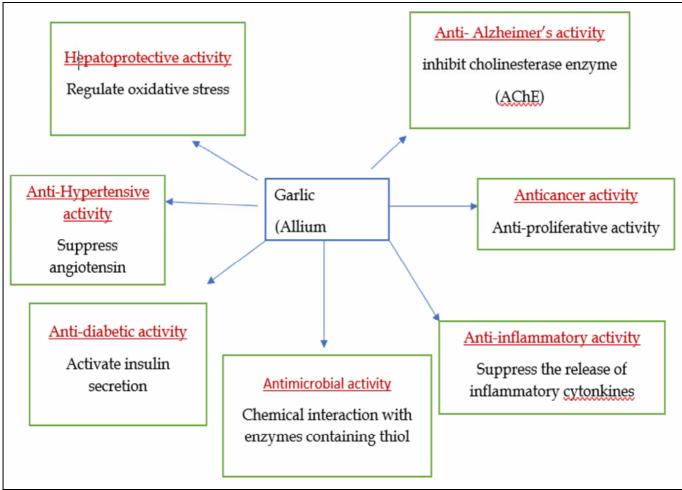
Properties of garlic

Organosulphur compounds	Beneficial effects	References
Garlic oil	Reduce toxicity of terbutylin	Liu and Xu (2007)
Aged garlic extract	Suppress the development of glycation end products and formation of glycation derived free radicals.	Ahmad and Ahmad (2006)
Aged game extract	Inhibit lipid peroxidation and oxidation of low-density lipoprotein (LDL)	Amagase <i>et al.</i> (2001) Lau(2006)
Aq. Garlic extract	Guard the tissue against nicotine induced oxidative disruption	Augusti(1996)
Allicin	Scavenges hydroxyl radicals	Prasad, Haxadaland Yu(1995)
Di-allyl tri-sulphate	Decrease the liver damage caused by carbon tetra chloride	Fukao <i>et al.</i> (2004) (S.G. Santhosha, Prakash Jamuna, S.N. Prabhavathin)

2,2-diphenyl-1-picrylhydrazyl (DDPH), a scavenging technique or method is used to identify the anti-oxidant property of raw garlic which shows a change in colour from violet to yellowish which indicates antioxidant property. However (HPLC) high performance liquid chromatography technique doesn't give appropriate conclusion about presence of allicin (Rahman, M. M *et al*).

Aged extracts which obtained from the leaves, they show best anti-oxidant property, however flower bulblets express less or no activity. (Anna Capasso)

Polyphenol groups helps to fight against the oxidative stress which affect membranes,nucleic acid, celluler component etc. (Florinda Fratianni



(2) Anti- microl	bial activity			
Product	Compound	Cell lines	Major effect	Reference
Garlic		Aspergillus versicolor	Suppress the growth of bacteria.	Fratianni, F.;
"Rosato"	Allicin	Penicillium expansum	Suppress the growth of bacteria.	Wallock-
And		Burkholderia cepacian	Suppress the growth of bacteria	Richards
"Caposele"		Staphylococcus aureus	Suppress the growth of bacteria	Guo, Y.J.
Aged garlic	Garlic oil	Candida albicans	Destruct the normal metabolism of bacteria.	Li, W.R.; Sh
extract		Penicillium		5. Li, W.R.; Shi
		funiculosum		
Raw garlic		15 patients with H.	Suppressing the Helicobacter pylori in the	Zardast, M.;
		pylori	stomach	Namakin
		infection		(Ao Shang)

(2) Anti- microbial activity

1.) Table showing Different garlic compounds showing acitivty against microorganisms e.g., *Penicillium expansum*, *Burkholderia cepacian*, *Staphylococcus aureus* etc.

Garlic is broad spectrum compound having antibacterial an anti-fungal property. (Fratianni, F.; Riccardi,). Different varieties of garlic "Rosato" and "Caposele" significantly inhibit the growth of microbes eg., Rosato inhibit Penicilliun citrinum and Aspergillus versicolor and Caposele have strong effect on inhibition of penicillium expansum (Fratianni, F.). Garlic oil also possess antibacterial properties to inhibit the growth of Staphylococcus aureus and Escherichia coli (Guo) (Ao Shang).

First man who purposed the attributes regarding antibacterial in case of garlic are Louis Pasteur in 1858 and exhibit the allicin as the main active constituent. (Block, 1992; Nagourney, 1998) (P. Avato', F. Tursi). Garlic contain 3 times sulphur compounds as similar as onions (Lawson) (N. Benkeblia). These compounds are high in reaction, volatile in nature, odour producing and lachrymatory (Block, Naganathan, Putman, & Zhao, 1992). (N. Benkeblia). The antibiotic activity of 1mg of llicin, which is S-methyl- Lcysteine has been equal to 15 IU of penicillin. s (Sivam, Lampe, Ulness, Swanzy, & Potter, 1997; Phayet al., 1999; Hsieh, Mau, & Huang, 2001;

Garlic oil also express antifungal properties by penetrating into cells and organelles, by disrupting the cell structure etc. Inhibit the groth of fungus e.g. Penicillium funiculosum etc. (Li, W.R.; Shi) (Ao Shang) Garlic extracts have shown strong activiy against fungus and suppress the synthesis of mycotoxins e.g. aflatoxin of Aspergillus parasiticus. (Lawson L.D.,) (Serge Ankri) Pure allicin have large amount of anti- candida property with Minimum MIC concentration of 7μ g/ml. (Hughes B.G.,) (Serge Ankri)

Anti-Fungal Activity test

- Anti-fungal activity test was performed under in vitro conditions, petri plate containing Yeast Glucose Chloramphenicol Agar) (Lattenzio, De Cicco, Di Venere) (N. Benkeblia)
- 2.) Essential oil extract was added to Yeast Glucose Chloramphenicol Agar
- 3.) For control test, add sterile water to YGCA media (50, 100, 200, 300 and 500 ml/l, vol/vol.).
- 4.) Inoculate fungi in above petri plates in centre of 5mm of diameter of mycelial mass of cultivated test fungi
- 5.) Cut with sterile cork borer
- 6.) Incubate petri plates at 21°c in dark
- 7.) Stop the incubation when mycelial growth mass of petri plates almost filled the petri dish
- Diameter was Identified by averaging the radial growth of mycelial mass in two orthogonal directions. (N. Benkeblia)

Use of raw garlic is to suppress the growth of Helicobacter pylori in stomach of patient to protect it from H. pylori from infection. Extract of garlic possess activity against H. pylori which is the main reason for gastric ulcers. (Zardast, M.; Namakin) (Ao Shang). Different garlic compounds and their preparations possess wide spectrum anti- bacterial properties against gram negative and grampositive bacteria involving Salmonella, Staphylococcus, Streptococcus etc. (Uchida Y., Takahashi T) (Serge Ankri)

In the recent studies it was observed that about 1 to 2 percent of garlic extract having anti-microbial activity. The main component of garlic S-methyl L-cysteine sulfoxide is enzymatically divided into methyl methanethiosulfinate, as same like allin is converted into allicin with the help of enzyme alliinase. MMTSO is a strong antimicrobial compound but as low as compared to allicin. (Small *et al.*, 1947). (Kyu hang kyung)

Product	Compound	Possible mechanism	Reference
Aged black garlic extract	Polyphenols	Stimulating the production of NO	Takashima, M.;
Aged Garlic	S-1-		Garcia-Villalon,
extract	propylenecysteine		A.L.;
Fermented garlic extract		Decreasing the expression of vascular endothelial cell adhesion molecule-1 and MMP9Increasing the expression of PKG and eNOS	Park, B.M.; Chun, H
Garlic	Alliin		Asdaq, S.M.; Inamdar, M.N
Raw garlic	Allyl methyl sulphide	Increasing Na+/K+-ATPase protein level	Khatua, T.N.; (Ao Shang).

(3) Prevention against cardiovascular disease

Compound	Dose	Effect	Reference
garlicin	64mg/day; drip	↓ Unstable angina	Li <i>et al</i> 2000
Garlic powder	300mg/day	↑ B.V. activity	Breithaupt-Grogler
Garlic powder	800mg/day	↓ plasma viscosity	Kiesewetter <i>et al</i> (Sanjay K Banerjee and Subir K Maulik)

In the following tables Compounds showing different properties against CVS.

Cardiovascular disease is directly linked to increase in various factors such as increase in platelet aggregation, increase in cholesterol, increased in low density lipoprotein, etc. (Rahman K). Different preparations e.g. oil of steam distilled garlic, garlic powder and aged garlic extract used in CVS diseases in clinical trials (Banerjee SK, Mukherjee PK, Maulik SK) (Khalid Rahman and Gordon M. Lowe).

(4) Effect of garlic products on platelet aggregation

(1) Effect of game produces on platelet aggregation			
Products	Duration	Effect	Reference
Garlic oil	In vitro	Suppress platelet aggregation	Vanderhock et al., 1980
Garlic powder	Four weeks	Suppress PA	Kiesewetter et al., 1991
Aged garlic extract	Ten months	Suppress PA	Sreiner and Lin, 1998
Dried garlic powder	Four weeks	No PA effect	Harenberg et al., 1988
Essential garlic oil	Ten days	No PA effect	Samson, 1982
Ether extract of garlic	Single dose	Increased coagulation time	Arora and Arora, 1981 (Sanjay K Banerjee and Subir K Maulik)

Table shows the activity of garlic products on platlet aggregation

In vitro clinical trials shows that garlic extract improves circulation of blood and blood properties. (Moriguchi T, *et al.*). Garlic improves the fluidity of erythrocytes isolated from hypecholesterolemic rats (Kempaiah RK, Srinivasan K) (Khalid Rahman and Gordon M. Lowe).

Garlic is major medicinal plant that is also used for the treatment of atherosclerosis and their prevention. Garlic based preparations are also used as anti-atherosclerotic effects such as lowering plasma fibrinogen level etc (Campbell JH, Efendy JL, Smith NJ, *et al*) (El-Sabban F, Abouazra H).

Anti- hyperlipidemic activity	Hypertensive effect	Anti platlet activity	Anti- atherosclerotic effect
Diallyl disulphide: Decrease in cholestrol synthesis, decrease in triglycerides and antioxidant.	Suppressing angiotensin converting enzyme (ACE), stimulation of nitric oxide and hydrogen sulphide.	Suppression of cyclooxygenase, reduce in synthesis pf thromboxane B2, leukotriene C4 and regulation of membrane phospholipase	Suppress cellular proliferative activity, synthesis of extracellular matrix and anti- inflammatory activity. (Igor A. Sobenin)

Different activities shown by garlic e.g., Antihypertensive, Anti-Hyperlipidaemic, Anti-Platelet activity, Anti-Atherosclerotic activity.

(5) Prevention against cancer

Garlic and its organosulphur compounds are highly effective against cancerous cells and to inhibit cancer

incidence in breast, colon, skin, uterine, oesophagus, and lung cancers (Amagase and Milner 1993) (J. A. Milner). Different mechanism of actions like suppression of Nnitroso compound (NOC) formation, enhanced DNA pair, decrease cell proliferation, and induction of apoptosis are there for prevention.

(6) Organosulphur compounds with antineoplastic properties

Compound	Cell type
Ajoene	Lymphocytes, leukemic, colonic
Diallyl sulphide	Leukocyte, prostate
S-allyl cysteine	Melanoma, neuroblastoma
Diallyl disulphide	Lung, colonic, skin, prostate
Allicin	Lymphoid (Sigounas et al. 1997). (J. A. Milner)

NOC's are biological and environmental indicator that may increase the chances of cancer in human body (Brown 1999; Ferguson 1999). Exposure of NOC may occur through the nasal route and through ingestion of the precursors (Lijinsky 1999) (J. A. Milner) Frequent study in china reported that consumption of allium vegetables or garlic compounds decreases the chances of stomach cancer (Adetumbi, M. A) (Martha Thomson). Allium vegetables include garlic, onion, leeks, garlic stalks are used to treat stomach cancer (Holly L. Nicastro).

(7) Inhibition of tumour growth

Acc. To Pan XY, Li FO, *et al.*, tested the effect of garlic extract and diallyl trisulphide on two human gastric cancer cell tissues. In this trial, garlic extract and allyl disulphide were more potent as compared to fluorouracil and mitomycin C to treat cancerous cells (Pan XY, Li FQ, Yu ILl, Xie GF, Wang H, Zliao LY, Zhaug O) (Benjamin H.S. Lau).

S-allyl mercaptocysteine is water soluble organo sulphur compound having anti-oxidant property and suppress the cell growth and increase the apoptosis in cancer cell lines (Xiao J, Ching YP) (Maryam Miraghajani). SAMC is preferable for longer term treatment because SAMC has very less side effects like irritating (Tanaka S, Haruma K) (Maryam Miraghajani) and having no adverse effects and interaction with medication of aged garlic. (Macan H,) (Maryam Miraghajani).

(8) Possible Mechanism

Major group which shows the anti-neoplastic or anticancer effect are one is lipid soluble and another one is water soluble.

Lipid soluble: - Allyl sulphur compounds

1.) diallyl trisulphide (DATS),

2.) diallyl disulphide (DADS)

Water soluble: - γ-glutamyl S-allyl cysteine groups

1.) S-allyl-cysteine (SAC)

2.) S-allylmercaptocysteine (SAMC) (Thomson and Ali, 2003) (S.H. Omar, N.A. Al-Wabel).

Mechanism include suppression of mutagenesis, inhibit tumour growth, modulation of enzyme activities, free radical scavenging. AGE (aged garlic extract) garlic directly act as to inhibit prostate cancer growth, inhibiting polyamines required for cell division and increase in breakdown of testosterone, all are needed for prostate cancer growth (Pinto *et al*).

S-allyl mercaptocysteine is required for inhibit the growth of breast cancer and also for erythroleukemia (Sigounas *et al.*,) (S.H. Omar, N.A. Al-Wabel).

Compound	Cancer type and mechanism	Reference
Ajoene	Leukaemia: Inhibit proliferation	Ahmed et al
	Induce apoptosis by suppressing the G2 phase of cell cycle.	
Allicin	Suppress the proliferation of cancer cells and increase apoptosis by	Oommen, Anto,
	activation of caspases.	Srinivas
S-allyl cysteine	Blockage of nitrosamine generation and bioactivation	Dion <i>et al.</i> (1997)
Diallyl di sulphide	Colon cancer: suppress the growth of neoplastic CMT-13 cells and	Chen, Chung,
	N acetyl transferase activity	Hsieh
Diallyl trisulfide	Prostate cancer: increase apoptosis by induction of protein bax and	Xiao <i>et al</i> .
	bak	(S.G. Santhosha)

(9) Compounds showing different cancer's mechanism

(10) Prevention against Alzheimer disease

Alzheimer is a form of dementia and amyloid beta(A β) deposition (Masters *et al.*, 1985) (Veer Bala Gupta). Alzheimer disease is directly related to psychiatric disease and multifactorial disease. Where almost many allopathic drugs failed to treat Alzheimer but nutraceuticals compounds, phytopharmaceuticals and ayurvedic medicines gave appropriate solution. AGE (aged garlic extract) and SAC (S-allyl cysteine) having anti amyloid property. These two compounds prevent hyperphosphorylation of tau, where tau is a protein helps the brain cells to work prop2.17. erly (Camila Calfio).

Garlic extract (GE) shows variety of properties and prevents cognitive decline by protecting neurons from A β neurotoxicity and apoptosis (Borek,2006) Different studies were done to show the positive effect of aqueous garlic extract and boiled garlic extract against the A β aggregation. GE and garlic constituents decrease A β neuronal apoptosis, by increasing the endogenous antioxidant defences (Peng *et al.*, 2002) (Veer Bala Gupta).

Frequent research proposed that beginning of $A\beta$ amyloid obtain from amyloidogenic processing of β amyloid precursor protein (APP) is the first episode in Alzheimer pathophysiology (Selkoe, 2000).

AGE is having multi phytochemicals like ajoene, polyphenols and also thiosulphates that possess multiple effects like anti apoptotic etc (Neelima B. Chauhan).

Age garlic extract shows neuroprotective effect as natural NASAID, natural anti apoptotic agent and to enhance memory having combination of single drug or ingredient now used for treatment of Alzheimer (Bordia A. 1978) (S M Shende).

Insulin secretagogue action of garlic compound increase the brain levels of insulin and insulin like growth factor (IGF) which reduce brain $A\beta$ burden and inhibit the activation of GSK β -3 and can potential Prevent tau phosphorylation(Ray B, Chauhan NB, Lahiri DK).

(11) Role of garlic in wound healing

Wound healing includes exchange between inflammatory cells and extracellular matrix throughout the three phases. These phases are inflammatory phase, proliferative phase and remodelling phase.

A.) During this phase damaged cells and pathogens are eliminate from the site of action.

B.) In proliferative phase, it includes the period of tissue growth that is started in reaction to stimulatory aspect startle assemble throughout the inflammatory phase. Angiogenesis is intermediate to wound healing and includes the generation of capillary blood vessels.

And lastly in remodelling period it includes the regeneration of extracellular matrix, formation of tissues etc.

Early wound healing dependent on angiogenesis and their presence is similar with granulation, migrating fibroblast and new collagen. Impaired angiogenesis ia a characteristic of chronic wound encountered with diabetes or artificial insufficiency (Li and Li, 2008; Folkman, 2003, 2007; Rees *et al.*, 1999; Bernstein *et al.*, 1994; Clark *et al.*, 2004; Yamaguchi and Yoshikawa, 2001; Tonnesen *et al.*, 2000; Kubo *et al.*, 2005) (Sohail Ejaz)

Wound healing is a physiological process and the inflammatory cells which increase the rate of healing are cytokines with the help of different pathway like decrease dehydration, increasing granulation tissue formation. Neutrophils release elatase and collagenase and tumour necrosis factor and macrophages appear at the wound site and differentiate into macrophages and secrete cytokines such as TGF- β and VEGF.

Macrophage involved in the phagocytosis and release PDGF, TGF- β , TNF- α , IL-1 AND IL-6 and they helps in the proliferative phase of wound healing. Lymphocytes are essential in the IL-2 formation which helps in recruit fibroblast.

Conventional aqueous antiseptics are not sustainable because the borders of chronic wounds are composed of sclerotic tissue impeding on effectual penetration of the products (Araújo LU; Dorai AA; Shah JM, Omar E; Lee YS, Wysocki A; Buganza Tepole A; Lademann O, Kramer A) Nader Pazyar

Burns are usually most noxious form of wound and minor burns are usually serve with topical ointments. Higher rate of mortality are because of burn wound difficulties especially to microbial infection. Many of the times wound healing drugs are not partially effective against microorganisms and are also not economically not available for all populations. Garlic has been used from thousands of centuries to treat infectious diseases and having various activities. Allium sativum is the major constituent which posses their activity against microorganisms and to heal wounds.

(Gould LJ., Fulton AT.; Church D, Elsayed S; Somboonwong J; Khashan AA. (Leila Ait Abderrahim)

Garlic possess antioxidant, antiplatelet, antifibrinolytic effect and having impact on microcirculation. Allium sativum consist large amount of alliin, allyl cysteine, allicin etc and these constituents are powerful antioxidants. In further studies it was observed that elevated antioxidants in wounds increase the healing of wounds. Apart from this, anti-inflammatory agents are also play major role in increasing healing of wound. Inflammatory cells like neutrophils, macrophages, lymphocytes etc they play important role to remove the infection by microorganisms and clear cellular debris (Sundaresan, Subramanian 2003) (Chung, 2006) (Rasik, Shukla, 2000; Farahpour *et al.*, 1015) (Guo, Dipietro) Mohammad Reza Farahpour

In an infected wound, pathogens easily exist in the protective Form which termed as biofilm. Biofilm is fixed in glycocalyx which is a mixture of bacterial secreted extracellular polysaccharide that make a safeguarding matrix adhering to the host surrounding tissue. Acc. To James *et al.* (2008) generation of polymicrobial biofilm in the chronic wounds such as foot ulcers, pressure ulcers etc.

In the previous studies the activity of Allium sativum and garlic extract (allicin) is to suppressing the generation of bacterial pathogens and allicin also been shown to elevate the susceptibility of P. aeruginosa to tobramycin and to phagocytosis. Garlic extract suppress the *P. aeruginosa* and *Acinetobacter baumanii* quorum sensing system they have been there to manage the biofilm enlargement.

In the present studies the activity of garlic ointment is suppressing biofilm growth by gram negative and grampositive bacteria in in-vitro wound biofilm model. Garlic ointment decrease the biofilm growth and development by *S. aureus, Staphylococcus epidermidis, P. aeruginosa* etc (Bjarnsholt *et al.*, 2008; Janes *et al.*, 2008) (Stoodley *et al.*, 2002; Sutherland, 2001) (James *et al.*, 2008) (Stoodley *et al.*, 2007) (Bjarnsholt *et al.*, 2005; Rasmussen *et al.*,) (Pushpalatha Nidadavolu).

Some herbs used in the treatment of cutaneous wounds

1.) Lemon	Lee YH, Chang J (140)
2.) Mimosa	Zippel J, Deters A (141)
3.) Soybean	Xu L, Choi TH, (142)
4.) Grape vine	Khanna S, Venojarvi M (143)
5.) Papaya	Nayak BS, Ramdeen R (144)
6.) Ginseng	Kimura Y, Sumiyoshi M (145)
7.) Olive oil	Kim HS, Noh SU (146)

(Nader Pazyar)

Conclusion

Wounds are biological process and in the end it causes scar formation. Various medicinal plants verified for healing wounds. Other than healing wounds, in severe conditions medicinal plants are highly effective. Garlic (*Allium sativum*) Family Liliaceae, a spice widely used all over the world having characteristic odor, produce highly effective action against cancer, act as anti-microbial, anti-fungal, Immune modulator, to prevent cardiovascular disorders. Garlic was also used for workers to increase their work capacity. In the present time, Garlic is still used worldwide as medicinal plant for treating variety of diseases and produce pharmacological effects. As summarized in this review, various properties of garlic. It is clear form the article that garlic in the diet should be considered to be important and mandatory to maintain good health.

Therefore, in the future more biological functions of garlic should be evaluated and more investigations should be conducted to deeply understand the mechanism of action of garlic.

References

- Abderrahim LA, Taïbi K, Abderrahim NA, Boussaid M, Rios-Navarro C, Ruiz-Saurí A. Euphorbia honey and garlic: Biological activity and burn wound recovery. Burns. 2019 Nov 1;45(7):1695-706.
- Ahmad, M. S., and Ahmed, N. (2006). Antiglycation property of aged garlic extract; possible role in prevention of diabetic complications. Journal of Nutrition, 136, 796s-799s.
- Ahmed, N., *et al.*, (2001). Ajoene a garlic derived natural compound enhances chemotherapy induced apoptosis in human Myeloid Leulaemia CD34- positiveresistant cells. Anticancer Research, 21, 3519-3524.
- Al-Waili, N. S., Saloom, K. Y., Akmal, M., Al-Waili, T. N., Al-Waili, A. N., Al-Waili, H., Ali, A. & Al-Sahlani, K.

(2007). Effects of heating, storage, and ultraviolet exposure on antimicrobial activity of garlic juice. J Med Food 10, 208–212

- Amagase, H. & Milner, J. A. (1993) Impact of various sources of garlic and their constituents on 7,12dimethylbenz(a)anthracene binding to mammary cell DNA. Carcinogenesis 14: 1627–1631.
- Amagase, H., *et al.*, (2001). Intake of garlic and its bioactive components. The Journal of Nutrition, 131, 955s-962s.
- Ankri S, Mirelman D. Antimicrobial properties of allicin from garlic. Microbes and infection. 1999 Feb 1;1(2):125-9.
- Araújo LU, Grabe-Guimarães A, Mosqueira VC, Carneiro CM, Silva-Barcellos NM: Profile of wound healing process induced by allantoin. Acta Cir Bras 2010; 25:460–466.
- Arora RC, Arora S: Comparative effects of clofibrate, garlic and onion on alimentary hyperlipemia. Atherosclerosis 1981, 39:447-452
- Asdaq, S.M.; Inamdar, M.N. Potential of garlic and its active constituent, S-allyl cysteine, as antihypertensive and cardioprotective in presence of captopril. Phytomedicine 2010, 17, 1016–1026.
- Augusti, K.T. (1996). Therapeutic values of garlic and onion. Indian Journal of Experimental Biology, 34, 634-640.
- Avato P, Tursi F, Vitali C, Miccolis V, Candido V. Allylsulfide constituents of garlic volatile oil as antimicrobial agents. Phytomedicine. 2000 Jun 1;7(3):239-43.
- BAIRY, K.L.; RAO, C.M. Wound healing profile of Ginko biloba. J. Nat. Remedies, v.1, p.25-27, 2001. CHUNG, L.Y. The antioxidant properties of garlic compounds: allyl cysteine, alliin, allicin, and allyl disulfide. J. Med. Food, v.9, p.205-213, 2006.
- Banerjee SK, Maulik SK. Effect of garlic on cardiovascular disorders: a review. Nutrition journal. 2002 Dec 1;1(1):4.
- Barku VYA, Boye A, Quansah N. Antioxidant and wound healing studies on the extracts of Corchorus olitorius leaf. World Essays J 2013;1:67–73
- Benjamin A. Lipsky and Christopher Hoey, Topical Antimicrobial Therapy for Treating Chronic Wounds: November 2009 – Volume 49 – Issue 10 – p 1541-1549
- Benkeblia N. Antimicrobial activity of essential oil extracts of various onions (*Allium cepa*) and garlic (*Allium sativum*). LWT-food science and technology. 2004 Mar 1;37(2):263-8.
- Bernstein, E.F., Chen, Y.Q., Tamai, K., Shepley, K.J., Resnik, K.S., Zhang, H., Tuan, R., Mauviel, A., Uitto, J. (1994). Enhanced elastin and fibrillin gene expression in chronically photodamaged skin. J Invest Dermatol 103:182–186
- Bjarnsholt, T., Jensen, P. Ø., Rasmussen, T. B., Christophersen, L., Calum, H., Hentzer, M., Hougen, H. P., Rygaard, J., Moser, C. & other authors (2005). Garlic blocks quorum sensing and promotes rapid clearing of pulmonary Pseudomonas aeruginosa infections. Microbiology 151, 3873–3880.
- Bjarnsholt, T., Kirketerp-Møller, K., Jensen, P. Ø., Madsen,K. G., Phipps, R., Krogfelt, K., Høiby, N. & Givskov,M. (2008). Why chronic wounds will not heal: a novel hypothesis. Wound Repair Regen 16, 2–10.
- Block, E., Naganathan, S., Putman, D., & Zhao, S. H. (1992). Allium chemistry: HPLC analysis of thiosulfonates

from onion, garlic, wild garlic, leek, scallion, shallot, elephant garlic, and Chinese chive. Uniquely high allyl to methyl ratios in some garlic samples. Journal of Agriculture and Food Chemistry, 40, 2418–2430.

- Block, E.: The organosulfur chemistry of the genus Allium Implications for the organic chemistry of sulfur. Angew. Chem. 31: 1135-1178, 1992.
- Borek C. 2006. Garlic reduces dementia and heart-disease risk. J Nutr 136: 810S–812S. Review
- Brown J. L. (1999) N-Nitrosamines. Occup. Med. 14: 839– 84
- Buganza Tepole A, Kuhl E: Systems-based approaches toward wound healing. Pediatr Res 2013; 73:553–563.
- Calfio C, Gonzalez A, Singh SK, Rojo LE, Maccioni RB. The Emerging Role of Nutraceuticals and Phytochemicals in the Prevention and Treatment of Alzheimer's Disease. Journal of Alzheimer's Disease. 2020 Jan 1(Preprint):1-9.
- Capasso A. Antioxidant action and therapeutic efficacy of *Allium sativum* L. Molecules. 2013 Jan;18(1):690-700.
- Chauhan NB. Effect of aged garlic extract on APP processing and tau phosphorylation in Alzheimer's transgenic model Tg2576. Journal of ethnopharmacology. 2006 Dec 6;108(3):385-94.
- Chen, G. W., Chung, J. G., Hsieh, C. L., & Lin, J. G. (1998). Effects of the garlic components diallyl sulfide and dially disulfide on Arylamine N-cetlytransferase activity in Human colon tumor cells, Food Chemistry and Toxicology, 36, 761-770.
- Church D, Elsayed S,Reid O,WinstonB, LindsayR. Burn wound infections.Clin Microbiol Rev 2006;19:403–34.
- Dion, M. E., Agler, M., & Milner, J. A. (1997). S-allyl cysteine inhibits nitrosomorpholine formation and bioactivation. Nutrition and Cancer, 28, 1-6.
- Diretto, G.; Rubio-Moraga, A.; Argandona, J.; Castillo, P.; Gomez-Gomez, L.; Ahrazem, O. Tissue-specific accumulation of sulfur compounds and saponins in different parts of garlic cloves from purple and white ecotypes. Molecules 2017, 22, 1359.
- Dorai AA: Wound care with traditional, complementary and alternative medicine. Indian J Plast Surg 2012; 45:418–424.
- Eckner, M. M.; Sticher, O.; Meier, B. Reversed-phase highperformance liquid chromatography of S-alk(en)yl-L-cysteine derivatives in *Allium sativum* including the determination of (+)- S-allyl-L-cysteine sulphoxide, γ-L-glutamyl-S-allyl-L-cysteine and γ-L-glutamyl-S-(trans-1-propenyl)-L-cysteine. J. Chromatogr. 1992, 625, 183-190
- Ejaz S, Chekarova I, Cho JW, Lee SY, Ashraf S, Lim CW. Effect of aged garlic extract on wound healing: a new frontier in wound management. Drug and chemical toxicology. 2009 Jul 1;32(3):191-203.
- El-Saber Batiha G, *et al.* Chemical constituents and pharmacological activities of garlic (*Allium sativum* L.): A review. Nutrients. 2020 Mar;12(3):872.
- Farahpour MR, Hesaraki S, Faraji D, Zeinalpour R, Aghaei M. Hydroethanolic *Allium sativum* extract accelerates excision wound healing: evidence for roles of mast-cell infiltration and intracytoplasmic carbohydrate ratio. Brazilian Journal of Pharmaceutical Sciences. 2017;53(1).
- Fratianni F, et al,. Phenolic constituents, antioxidant, antimicrobial and anti-proliferative activities of

different endemic Italian varieties of garlic (*Allium sativum* L.). Journal of Functional Foods. 2016 Mar 1; 21: 240-8.

- Fratianni, F.; Riccardi, R.; Spigno, P.; Ombra, M.N.; Cozzolino, A.; Tremonte, P.; Coppola, R.; Nazzaro, F. Biochemical characterization and antimicrobial and antifungal activity of two endemic varieties of garlic (*Allium sativum* L.) of the campania region, southern Italy. J. Med. Food. 2016, 19, 686–691
- Fukao, T., *et al.*, The effect of allyl sulphides on the induction of phase 2 detoxification enzymes and liver injury by carbon tetrachloride. Food Chemistry and Toxicology, 42, 743.
- Garcia-Villalon, A.L.; Amor, S.; Monge, L.; Fernandez, N.; Prodanov, M.; Munoz, M.; Inarejos-Garcia, A.M.; Granado, M. In vitro studies of an aged black garlic extract enriched in S-allylcysteine and polyphenols with cardioprotective effects. J. Funct. Foods 2016, 27, 189– 200.
- Gould LJ, Fulton AT. Wound healing in older adults. R I Med J 2013; 99:34–6,
- GUO, S.; DIPIETRO, L.A. Factors affecting wound healing. J. Dent. Res., v.89, n.3, p.219-229, 2010.
- Guo, Y.J. Experimental study on the optimization of extraction process of garlic oil and its antibacterial effects. Afr. J. Tradit. Complement. Altern. Med. 2014, 11, 411–414. [
- Guo, Y.J.; Deng, G.F.; Xu, X.R.; Wu, S.; Li, S.; Xia, E.Q.; Li, F.; Chen, F.; Ling, W.H.; Li, H.B. Antioxidant capacities, phenolic compounds and polysaccharide contents of 49 edible macro-fungi. Food Funct. 2012, 3, 1195–1205.
- Gupta VB, Indi SS, Rao KS. Garlic extract exhibits antiamyloidogenic activity on amyloid beta fibrillogenesis: relevance to Alzheimer's disease. Phytotherapy Research: An International Journal Devoted to Pharmacological and Toxicological Evaluation of Natural Product Derivatives. 2009 Jan;23(1):111-5.
- Harenberg J, Giese C, Zimmermann R: Effect of dried garlic on blood coagulation, fibrinolysis, platelet aggregation and serum cholesterol levels in patients with hyperlipoproteinemia. Atherosclerosis 1988, 74:247-249
- Hermans, M.H.E. and T. Treadwell, An Introduction to Wounds, in Microbiology of Wounds, S. Percival and K. Cutting, Editors. 2010, CRC Press: Boca Raton, Fla. p. 83-134.
- Hughes B.G., Lawson L.D., Antimicrobial effects of Allium sativum L. (garlic) Alliumampeloprasum (elephant garlic) and Allium cepa L. (onion) garlic compounds and commercial garlic supplement products, Phytother. Res. 5 (1991) 154–158.
- James, G. A., Swogger, E., Wolcott, R., Pulcini, E., Secor, P., Sestrich, J., Costerton, J. W. & Stewart, P. S. (2008). Biofilms in chronic wounds. Wound Repair Regen 16, 37–44.
- Kempaiah RK, Srinivasan K. Influence of dietary spices on the fluidity of erythrocytes in hypercholesterolaemic rats. Br J Nutr. 2005; 93:81–91
- Keusgen, M.; Junger, M.; Krest, I.; Schoning, M. Development of a biosensor specific for cysteine sulfoxides. J. Biosens. Bioelectron. 2003, 18, 805-812.

- Khanna S, Venojarvi M, Roy S, Sharma N, Trikha P, Bagchi D, Bagchi M, Sen CK: Dermal wound healing properties of redox-active grape seed proanthocyanidins. Free Radic Biol Med 2002; 33:1089–1096
- Khashan AA. Antibacterial activity of garlic extract (*Allium sativum*) against Staphylococcus aureus in vitro. GJBB 2014;3: 346–348.
- Khatua, T.N.; Borkar, R.M.; Mohammed, S.A.; Dinda, A.K.; Srinivas, R.; Banerjee, S.K. Novel sulfur metabolites of garlic attenuate cardiac hypertrophy and remodeling through induction of Na+/K +-ATPase expression. Front. Pharmacol. 2017, 8, 18
- Kiesewetter H, Jung F, Pindur G, Jung EM, Mrowietz C, Wenzel E: Effect of garlic on thrombocyte aggregation, microcirculation, and other risk factors. Int J Clin Pharmacol Ther Toxicol 1991, 29:151-155
- Kubec, R.; Svobodova, M.; Velisek, J. Gas chromatographic determination of S-alk(en)ylcysteine sulfoxides. J. Chromatogr. A 1999, 862, 85-94.
- Kubo, S., Inui, T., Hasegawa, H., Yoshimine, T. (2005). Repair of intractable cerebrospinal fluid rhinorrhea with mucosal flaps and recombinant human basic fibroblast growth factor: technical case report. Neurosurgery 56: E627.
- KYUNG KH, PARK KS, KIM YS. Isolation and characterization of bacteria resistant to the antimicrobial activity of garlic. Journal of food science. 1996 Jan;61(1):226-9.
- Lademann O, Kramer A, Richter H, Patzelt A, Meinke MC, Czaika V, Weltmann KD, Hartmann B, Koch S: Skin disinfection by plasmatissue interaction: comparison of the effectivity of tissue-tolerable plasma and a standard antiseptic. Skin Pharmacol Physiol 2011;24: 284–288
- Lattenzio, V., De Cicco, V., Di Venere, D., Lima, G., & Salerno, M. (1994). Antifungal activity of phenolics against fungi commonly encountered during storage. Italian Journal of Food Science, 1, 23–30.
- Lau BH, Tadi PP, Tosk JM. *Allium sativum* (garlic) and cancer prevention. Nutritionresearch. 1990 Aug 1;10(8):937-48.
- Lawson L.D., The composition and chemistry of garlic cloves and processed garlic. in: Koch H.P., Lawson L.D. (Eds.), Garlic: the science and therapeutic application of *Allium sativum* L., Williams and Wilkins, Baltimore, 1996, pp. 37–108.
- Lawson, L. D. (1996). Garlic: a review of its medicinal effects and indicated active compounds. In L. D. Lawson, & R. Bauer (Eds.), Phytomedicines of Europe: Their chemistry and biological activity (pp. 176–209). Washington DC: ASC Press
- Lee YH, Chang JJ, Chien CT, Yang MC, Chien HF: Antioxidant sol-gel improves cutaneous wound healing in streptozotocin-induced diabetic rats. Exp Diabetes Res 2012; 2012: 504693.
- Lee YS, Wysocki A, Warburton D, Tuan TL: Wound healing in development. Birth Defects Res C Embryo Today 2012; 96:213–222.
- Li, V.W., Li, W.W. (2008). Antiangiogenesis in the treatment of skin cancer. J Drugs Dermatol 7:s17–s24.
- Li, W.R.; Shi, Q.S.; Dai, H.Q.; Liang, Q.; Xie, X.B.; Huang, X.M.; Zhao, G.Z.; Zhang, L.X.Antifungal activity, kinetics and molecular mechanism of action of garlic oil against Candida albicans. Sci. Rep. 2016, 6, 22805

- Li, W.R.; Shi, Q.S.; Liang, Q.; Huang, X.M.; Chen, Y.B. Antifungal effect and mechanism of garlic oil on penicillium funiculosum. Appl. Microbiol. Biot. 2014, 98, 8337–8346.
- Lijinsky, W. (1999) N-Nitroso compounds in the diet. Mutat. Res. 443: 129–138
- Liu, J.; Ji, F.; Chen, F.M.; Guo, W.; Yang, M.L.; Huang, S.X.; Zhang, F.; Liu, Y.S. Determination of garlic phenolic compounds using supercritical fluid extraction coupled to supercritical fluid chromatography/tandem mass spectrometry. J. Pharm. Biomed. Anal. 2018, 159, 513–523.
- Liu,H. G., and Xu, L.H.(2007). Garlic oil prevents tributylin-induced oxidative damage in vivo and in vitro. Journal of Food Protection, 70, 716-721.
- Macan H, Uykimpang R, Alconcel M, Takasu J, Razon R, Amagase H, *et al.* Aged garlic extract may be safe for patients on warfarin therapy. J Nutr 2006; 136:793S-795S.
- Masters CL, Simms G, Weinman N, Multhaup G, McDonald BL, Beyreuther K. 1985. Amyloid plaque core protein in Alzheimer disease and Down syndrome. Proc Natl Acad Sci USA 82: 4245–4249
- Milner JA. A historical perspective on garlic and cancer. The Journal of nutrition. 2001 Mar;131(3):1027S-31S.
- Miraghajani M, Rafie N, Hajianfar H, Larijani B, Azadbakht L. Aged garlic and cancer: A systematic review. International journal of preventive medicine. 2018;9.
- Moore, K., Cell Biology of Normal and Impaired Healing, in Microbiology of Wounds, S. Percival and K. Cutting, Editors. 2010, CRC Press: Boca Raton, Fla. p. 151-186.
- Moriguchi T, Takasugi N, Itakura Y. The effects of aged garlic extract on lipid peroxidation and the deformability of erythrocytes. J Nutr. 2001;131: 1016S– 9S.
- Nayak BS, Ramdeen R, Adogwa A, Ramsubhag A, Marshall JR: Wound-healing potential of an ethanol extract of Carica papaya (Caricaceae) seeds. Int Wound J 2012; 9:650–655
- Nicastro HL, Ross SA, Milner JA. Garlic and onions: their cancer prevention properties. Cancer prevention research. 2015 Mar 1;8(3):181-9.
- Nidadavolu P, Amor W, Tran PL, Dertien J, Colmer-Hamood JA, Hamood AN. Garlic ointment inhibits biofilm formation by bacterial pathogens from burn wounds. Journal of medical microbiology. 2012 May 1;61(5):662-71.
- Omar SH, Al-Wabel NA. Organosulfur compounds and possible mechanism of garlic in cancer. Saudi Pharmaceutical Journal. 2010 Jan 1;18(1):51-8.
- Oommen, S., *ET AL*.,(2006). Effect of garlic on isoniazid and rifampicin-induced hepatic injury in rats.World Journal of Gastroenterology, 12, 636-639.
- O'Toole, E.A. and J.E. Mellerio, Wound Healing, in Rook's Textbook of Dermatology, T. Burns, *et al.*, Editors. 2010, Wiley-Blackwell: Oxford, UK. p. 14.1-14.27. 14. Moore, K., Cell Biology of Normal and Impaired Healing, in Microbiology.
- Pan XY, Li FQ, Yu ILl, Xie GF, Wang H, Zliao LY, Zhaug O. Experimental chemotherapy of human gastric cancer cell lines in vitro and in nude mice. Chinese J Oncology 10:15-18, 1988
- Park, B.M.; Chun, H.; Chae, S.W.; Kim, S.H. Fermented garlic extract ameliorates monocrotaline-induced

pulmonary hypertension in rats. J. Funct. Foods 2017, 30, 247–253.

- Pazyar N, Yaghoobi R, Rafiee E, Mehrabian A, Feily A. Skin wound healing and phytomedicine: a review. Skin pharmacology and physiology. 2014;27(6):303-10.
- Pazyar N, Yaghoobi R: Ginseng: potential for the antileishmanial arsenal? Sao Paulo Med J 2013; 131:279–280Kim HS, Noh SU, Han YW, Kim KM, Kang H, Kim HO, Park YM: Therapeutic effects of topical application of ozone on acute cutaneous wound healing. J Korean Med Sci 2009; 24:368–374.
- Peng Q, Buz'Zard AR, Lau BH. 2002. Neuroprotective effect of garlic compounds in amyloid-beta peptide-induced apoptosis in vitro. Med Sci Monit 8: BR328–BR337.
- Pinto, J.T., Qiago, C., Xing, J., Suffoletto, B.P., Schubert, K.B., Rivlin, R.S., Huryk, R.F., Bacich, D.J., Heston, W.D., 2000. Alteration of prostate biomarker expression and testosterone utilization in human LNCaP prostate carcinoma cells by garlic derived Sallylmercaptocysteine. Prostate 45, 304–314.
- Prasad, K., Haxdal, A., and Yu, M. (1995). Antioxidant property of Allicin, an active principle in garlic. Molecular and Cellular Biochemistry, 148, 183-189.
- Rahman K, Lowe GM. Garlic and cardiovascular disease: a critical review. The Journal of nutrition. 2006 Mar 1;136(3):736S-40S.
- Rahman MM, Fazlic V, Saad NW. Antioxidant properties of raw garlic (*Allium sativum*) extract. International Food Research Journal. 2012 Aug 1;19(2):589-91.
- RASIK, A.M.; SHUKLA, A. Antioxidant status in delayed healing type of wounds. Int. J. Exp. Pathol., v.81, n.4, p.257-263, 2000.
- Ray B, B Chauhan N, K Lahiri D. The "Aged Garlic Extract"(AGE) and One of its Active Ingredients S-Allyl-LCysteine (SAC) as Potential Preventive and Therapeutic Agents for Alzheimer's Disease (AD). Current medicinal chemistry. 2011 Aug 1;18(22):3306-13.
- Samson RR: Effects of dietary garlic and temporal drift on platelet aggregation. Atherosclerosis 1982, 44:119-120
- Santhosha S.G., *et al.*, Bioactive components of garlic and their physiological role in health maintenance: A review: September 2013 vol 3 pg. no. 59-74.
- Santhosha SG, Jamuna P, Prabhavathi SN. Bioactive components of garlic and their physiological role in health maintenance: A review. Food Bioscience. 2013 Sep 1;3:59-74.
- Selkoe, D.J., 2000. Toward a comprehensive theory for Alzheimer's disease Hypothesis: Alzheimer's disease is caused by the cerebral accumulation and cytotoxicity of amyloid beta-protein. Annals of New York Academy of Sciences 924, 17–25.
- Shah JM, Omar E, Pai DR, Sood S: Cellular events and biomarkers of wound healing. Indian J Plast Surg 2012; 45:220–228.
- Shang, A.; Cao, S.-Y.; Xu, X.-Y.; Gan, R.-Y.; Tang, G.-Y.; Corke, H.; Mavumengwana, V.; Li, H.-B. Bioactive Compounds and Biological Functions of Garlic (*Allium sativum* L.). *Foods* July 2019, 8, 7, 246.
- Sigounas, G., Hooker, J., Anagnostou, A. & Steiner, M. (1997) S-Allylmercaptocysteine inhibits cell proliferation and reduces the viability of erythroleukemia, breast, and prostate cancer cell lines. Nutr. Cancer 27: 186–191.

- Sigounas, G., Hooker, J., Angnostou, A., Steiner, M., 1997. S-allyl mercaptocysteine inhibits cell proliferation and reduces the viability of erythroleukemia, breast and prostate cancer cell lines. Nutr. Cancer 27, 186–191
- Sivam, G. P., Lampe, J. W., Ulness, B., Swanzy, S. R., & Potter, J. D. (1997). Helicobacter pylori—in vitro susceptibilityto garlic (*Allium sativum*) extract. Nutrition and Cancerology, 27, 118–121.
- Sobenin IA, Myasoedova VA, Iltchuk MI, Zhang DW, Orekhov AN. Therapeutic effects of garlic in cardiovascular atherosclerotic disease. Chinese journal of natural medicines. 2019 Oct 1;17(10):721-8.
- Somboonwong J, Kankaisre M, Tantisira B, Tantisira MH. Wound healing activities of different extracts of Centella asiatica in incision and burn wound models: an experimental animal study. BMC Complement Altern Med 2012; 12:1, doi: http://dx.doi.org/10.1186/1472-6882-12-103.
- Steiner M, Lin RS: Changes in platelet function and susceptibility of lipoproteins to oxidation associated with administration of aged garlic extract. J Cardiovasc Pharmacol 1998, 31:904- 908
- Stoodley, P., Sauer, K., Davies, D. G. & Costerton, J. W. (2002). Biofilms as complex differentiated communities. Annu Rev Microbiol 56, 187–209
- Sundaresan, S.; Subramanian, P. Garlic modulates lipid peroxidation and antioxidant status during N-Nitrosoethylamine-induced hepatic tumorigenesis. Plant Foods Hum. Nutr., n.58, n.3, p.1-8, 2003.
- Szychowski, K.A.; Rybczynska-Tkaczyk, K.; Gawel-Beben, K.; Swieca, M.; Karas, M.; Jakubczyk, A.; Matysiak, M.; Binduga, U.E.; Gminski, J. Characterization of active compounds of different garlic (*Allium sativum* L.) cultivars. Pol. J. Food Nutr. Sci. 2018, 68, 73–81.
- Tanaka S, Haruma K, Yoshihara M, Kajiyama G, Kira K, Amagase H, *et al.* Aged garlic extract has potential suppressive effect on colorectal adenomas in humans. J Nutr 2006;136: 821S-6S.
- Thomas Hess, Cathy BSN, RN, CWOCN Checklist for Factors Affecting Wound Healing, Advances in Skin & Wound Care: April 2011 - Volume 24 - Issue 4 - p 192.
- Thomson M, Ali M. Garlic [*Allium sativum*]: a review of its potential use as an anti-cancer agent. Current cancer drug targets. 2003 Feb 1;3(1):67-81.
- Thomson, M., Ali, M., 2003. Garlic [*Allium sativum*]: a review of its potential use as an anti-cancer agent. Curr. Cancer Drug. Target. 3, 67–81.

- Tonnesen, M.G., Feng, X., Clark, R.A. (2000). Angiogenesis in wound healing. J Invest Dermatol Symp Proc 5:40– 46.
- Uchida Y., Takahashi T., Sato N., The characteristics of the antibacterial activity of garlic, Jpn J. Antibiotics 28 (1975) 638–642.
- Ushijima, M.; Takashima, M.; Kunimura, K.; Kodera, Y.; Morihara, N.; Tamura, K. Effects of S-1propenylcysteine, a sulfur compound in aged garlic extract, on blood pressure and peripheral circulation in spontaneously hypertensive rats. J. Pharm. Pharmacol. 2018, 70, 559–565.
- Vanderhock JY, Makheja AN, Bailey JM: Inhibition of fatty acid oxygenases by onion and garlic acts. Evidence for the mechanism by which these oils inhibit platelet aggregation. Biochem Pharmacol 1980, 29:3169-3173
- WALKER, H.L.; MASON Jr., A.D. A standard animal burn. J. Trauma, v.8, n.6, p.1049-1051, 1968.
- Wallock-Richards, D.; Doherty, C.J.; Doherty, L.; Clarke, D.J.; Place, M.; Govan, J.R.W.; Campopiano, D.J. Garlic revisited: Antimicrobial activity of allicincontaining garlic extracts against Burkholderia cepacia complex. PLoS ONE 2014, 9, e112726.
- Xiao J, Ching YP, Liong EC, Nanji AA, Fung ML, Tipoe GL, *et al.* Garlic-derived S-allylmercaptocysteine is a hepato-protective agent in non-alcoholic fatty liver disease in vivo animal model. Eur J Nutr 2013; 52:179-91
- Xiao, D., Choi, S., Johnson, D. E., Vogel, V. G., Johnson, C. S., Trump, D. L., *et al.*, (2004). Diallyl trisulfideinduced apoptosis in human prostate cancer cells involves c-Jun N-terminal kinase and extracellularsignal regulated kinase-mediated phosphorylation pf Bcl-2. Oncogene, 23, 5594-5606.
- Xu L, Choi TH, Kim S, Kim SH, Chang HW, Choe M, Kwon SY, Hur JA, Shin SC, Chung JI, Kang D, Zhang D: Anthocyanins from black soybean seed coat enhance wound healing. Ann Plast Surg 2013; 71:415–420.
- Zardast, M.; Namakin, K.; Kaho, J.E.; Hashemi, S.S. Assessment of antibacterial effect of garlic in patients infected with Helicobacter pylori using urease breath test. Avicenna J. Phytomed. 2016, 6, 495–501
- Zippel J, Deters A, Hensel A: Arabinogalactans from Mimosa tenuiflora (Willd.) Poiret bark as active principles for wound-healing properties: specific enhancement of dermal fibroblast activity and minor influence on HaCaT keratinocytes. J Ethnopharmacol 2009; 124:391–396