

Usage of Evidence-Based Plant Medicine in Multimorbidity Lymphadenopathy Conditions

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Abstract

Immunoglobulin therapy is a popular trend and play a probability in assessing patient having defects such as Ig G diseases such as chronic lymphocytic leukemia, multiple myeloma, hypoimmune globulinaemia, and Ig M diseases such as hyperimmune globulinaemia, hepatitis, Ebola virus related with hemorrhagic fever, hepatitis B, Rabies, Tetanus toxin. This indispensable therapy helps to resolve various hypothetical disease conditions such as sepsis such as multiple sclerosis, neonatal sepsis, pediatric HIV/AIDS, PANDAS (pediatric autoimmune neuropsychiatric condition), a disorder associated with streptococcal cystic fibrosis), and unpredictable factor Chronic fatigue syndrome. This therapy involves maintaining a positive relationship with stress management with various factors such as innovation-based learning, Career concern, role ambiguity, rotating work shifts, role conflict, occupational demands, lack of participation in decision making, work overload, work under load, poor working conditions, lack of group cohesiveness, interpersonal and intergroup conflict, organizational changes and lack of social support. This article seeks to explicate and contextualize the empirical relationship between rigid chromophoric group plant constituents as well as shifting coincidence with various auxochromic groups which helps to reduce or increase or balancing about immunochemical reaction as well as management of lymphadenopathy conditions.

Keywords: IgM, IgG, Multimorbidity, Herbal medicines, Lymphadenopathy.

1. Introduction

According to EBM (Evidence-based medicines) treatment, scientists (author) focus on issues to prioritize therapeutic goals, selection of treatments based on patient values and preferences, and seek help to assist patients in adhering to agreed therapeutic regimens. Results from our research will be disseminated via publication in peer-reviewed journals. The impact of choosing evidence-based comorbidity treatment is creating tremendous benefits associated with heart diseases, hypertension, respiratory diseases, and mental health problems including dementia, cerebrovascular diseases, joint diseases, diabetes, and sensory impairment.

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This is an evidence map denoting about knowledge cluster of end-of-life care preferences of patients with multimorbidity. The mixed-methods appraisal tool is helping for multiple complications of patients on the basic selection of different plants having suppressive agents, plants having oxidate burst activity, plants having macrophage migration inhibitory factor, plants selected to manage an autoimmune disease, plants selected to rectify ADR factor, plants selected to block macrophage inhibiting factor and plant list which helps to involve in lymphatic drainage activity. Circular Queue linear data predict Isotope switching (Immunoglobulin class switching) in change over from one class immunoglobulin to another class immunoglobulin during situation incidence based on FIFO analysis EBPM treatment as per clinical practice guidelines.

Spontaneous stress is triggered by the current immunization COVID 19 vaccine, infections, emotional stress, and physical stress [1]. Fortunately, it is balanced with Non-pharmacological treatment such as Tai chi significant in the healing process, and Tai chi Qigong meditation motion.

2. Evidence-Based Plant Medicine Analysis for Multiple Complications

2. 1. Selection of plant list for multiple complications

Multiple complications usually involve an aggravation worsening in severity of the disease or the advances in the development of new signs, symptoms, or pathological changes which may become widespread throughout the body and affect other organ systems. The development of complications depends on several factors including the degree of vulnerability, health status, susceptibility, age, and immune system condition.

Complications in medicine are unfavorable results of disease, health condition, or treatment. To prevent common illnesses and complications such as Iatrogenic reaction complications (Medication, surgical, diagnostic, HAI healthcare-associated infections), Cardiovascular complications (Atrial fibrillation), Metabolic complications (Diabetes mellitus, Diabetes with CVS, Diabetic neuropathy, Foot damage), Neurological/Psychiatric complications (Hepatic encephalopathy, seasonal affective disorders [2]. Paradoxical reaction), Reproductive complications (Pregnancy-related hypertension, Gestational diabetes, preterm labor, miscarriage, stillbirth, Hyperemesis gravidarum) Respiratory complications (Scarlet fever, Rheumatic fever, meningitis, Glomerulonephritis, otitis media, Toxic shock syndrome) and surgical procedure complications (Puerperal fever, erectile dysfunction, Urinary incontinence, malignant hyperthermia)

To rectify multiple complications of diseases, follow up medical procedures via different usage of plant selection. The reason behind choosing plant selection will depend upon disease spreadability, Geographical predisposition, affected race, Genetic mutation in humans as well as a microorganism, Serotaxonomical classification-antigen antibody reaction, and development of abnormal plasma cell modification. On basis of Toxicity profile data concerning idiopathic and self-limiting -Lymphadenopathy is a major causative disease of lymph nodes in which they are abnormally deviating from average size or consistency [3]. Acute toxicity -Lymphadenopathy express the condition as a bacterial or viral disease [4]. Chronic toxicity Lymphadenopathy expresses the condition as Tuberculous lymphadenopathy and cat-scratch disease. Adverse drug reaction idiopathic self-limiting -Lymphadenopathy express the condition as HIV/AIDS,

autoimmune disease cancer. Inflammatory Lymphadenopathy expresses condition as lymphadenitis, scrofula, cervical lymphadenitis, Lymphadenitis associated with tuberculosis, and Lymphadenitis associated with Non-tuberculosis)

Acute infectious lymphadenopathy expresses the condition as a reduced decremental count of CD4+T cells and NK cells. Chronic infectious Lymphadenopathy express condition as NK/T cells lymphoma-cytotoxic syndrome. As per trending aspects in medical treatment -allopathic medicine commonly used drugs are clindamycin, Trimethoprim, amoxicillin, and ciprofloxacin in various infectious diseases such as Staphylococcal, streptococcal, H.Pylori, E.Coli, H.influenzae, mycobacterial infectious)Unfortunately, the human system could not able to adapt medications due to antibiotic resistance using the usage of antibiotics that could affect genetic mutation in the human genome. Hypothetical phenomena for using antibiotics in the treatment of lymphadenitis attributed to taking antibacterial medications which reflected the person is affected by bacterial infection, not viral infections. Swelling of lymph nodes cannot be controlled by antibacterial drugs which reflected the person is affected by a viral infection, not bacteria. Both the treatment ineffective condition, the blood is poisoned with an accumulation of endotoxin which could cause detrimentally synergistic effect on the congested lymphatic system. This condition will tend to make humans multiple complications-systemic infectious personalities as well as immunocompromised patients.

The human body cannot adapt to the situation -antibacterial, antiviral, toxin removal treatment, human is affected by pyogenic abscess, pyaemic abscess, and cold abscess [5]. On pathogenicity factor, an abscess form as an outcome of bacterial infection although a fungal or protozoal infection can also trigger an abscess formation. The internal abscess [6] can be identified due to weight loss and mental confusion.

On dissemination disease, spreadability factor related with lymphatic fluid blood or sepsis condition cannot reconstruct any draining lymphatic treatment the person can be fatal. From this point of view, multiple complications cannot be reorganized and attributed to co-morbidities.

Based on mutagenicity factor human-infected human, genetic variant human -genetic variant microorganisms, infected human-infected human factorial design will result in Zellweger syndrome -cerebrohepatorenal syndrome. This Usher syndrome [7] is caused due to autosomal recessive pattern -thickening and expansion of congenital disorders which are characterized by depletion or absence of functional peroxisomes in the cells of an individual. This is a genetic disorder leukodystrophy passed from parents to children that can be identified as a primer mover concept ATP -Dependent process (cellular function) altered and newborn can bring about brain, liver and kidney problems as well as difficulty feeding or moving and can put at risk in nerves and metabolism. Because of additive property enhancement in drug-addicted patients taking mephedrone -4 methyl methcathinone [8], [9] can bring about hyperactivity in CNS stimulants prolonged usage of medications can imperil facial blindness. (Red-green color blindness)

Mutation in melanopsin (opsin 4)-intrinsically photosensitive retinal ganglion cells can disrupt the mechanism of cyclic AMP-dependent process which could produce seasonal affective disorders as well as prosopagnosia (Red-green color blindness is followed by blue, yellow color blindness). In combination with viral vectors such as adenovirus -opsin 4 green fluorescent protein can occasionally bring about deaf-blindness, especially in sexually transmitted diseases. Lack

of serotonin, insufficient melatonin, a serotonin-derived hormone, Improper activation of G-Protein coupled receptor, mutational genetic defects in chromosome 10 Opn4 [10](Human melanopsin gene), Improper Third eye-opener development -pineal gland, a missense mutation in opn4, PIOL due to invading microorganism genetic variant and malnutrition endangering effect Nyctalopia (adaptation to darkness) and loss of peripheral vision -Tunnel vision are the factors attributed to facial blindness.

Impairment of motor dysfunction can produce neurological manifestations such as meningitis [11] or encephalitis [12], especially it alters or reduces consciousness, headache, fever, confusion, and stiff neck [13], [14], [15], [16], and vomiting. Complications in encephalitis will lead to enhance systematic infections such as seizures, Hallucinations, aphasia, memory problems, and conductive hearing loss which lead to endangering effects of autoimmune diseases such as HSV, Rabies, bacteria, fungi, or parasites. In extreme conditions related to a hypersensitivity reaction in association with left ventricular dysfunction and ECG changes. Hypothetical relationship to increase C-reactive protein (Cysteine-rich protein) can put in jeopardy effect on myocarditis (inflammation of heart muscle) due to viral infection [17]. Different types of plants as immunosuppressive agents are shown in Table 1.

Table 1. Denote Different Types of Plants (Immunosuppressive Agents), Chemical Constituents, Mechanism of Action, and Application.

S.no	LIST OF PLANTS	TAXONOMICAL CLASSIFICATION (Family, Class, Phylum)	CHEMICAL CONSTITUENTS	MECHANISM OF ACTION & APPLICATION
1.	Astragalus membranous [18], [19]	Fabaceae Eudicots Tracheophytes	Polysaccharides, Saponins, Flavonoids, Triterpenoid saponins	Downregulate Expression of phosphorylated epidermal growth factor & phosphorylated -Akt protein & upregulate expression of p-JNK. <u>APPLICATION</u> Antiaging Protect skin fibroblasts
2.	Ganoderma Lucidum [20]	Ganodermataceae Agaricomycetes Basidiomycota	Ganoderic acid A,B,E,CI,F Ganolucidic acid, Ganoderiol, Ganoderma triol, lucidumol	Downregulate Metastasis & regulate insulin resistance <u>APPLICATION</u> Anti-HIV AND Anti complement Activity
3.	Calendula officinalis [80]	Asteraceae Eudicots Tracheophytes	Zeaxanthin, Flavoxanthin, Auroxanthin, Carotenoids, Triterpenoid esters, Lutein,	Inhibit replication of HIV-1 in acutely infected lymphocytic MOLT-4 Cells. And PHA stimulated peripheral blood mononuclear cells <u>APPLICATION</u>

			Sesquiterpene glycosides	Anti-HIV activity and antimutagenic activity
4.	Goldenseal Hydrastis [88], [89]	Ranunculaceae Eudicots Tracheophytes	Isoquinoline alkaloids, Hydrastine, Berberine, berberastine, hydrastinine, canadine, canalidine	Regulate oxidative Glycolytic pathway <u>APPLICATION:</u> Treatment of Gonococcal ophthalmia, Actinic conjunctivitis, Photophthalmia
5.	Lonicera japonica [25], [26]	Caprifoliaceae, Eudicots, Tracheophytes	Loniceracetalides A&B Saponins loniceramide A & B Loniceramide C Japopenoid D&E, Ioniflavone	Regulate the Glycogenolysis process. <u>APPLICATION</u> Treatment for Hepatic steatosis and insulin resistance Anti HBV, Antihepatoma, HINI influenza A virus.
6.	Elderberry Sambucus Callicarpa [27], [28]	Caprifoliaceae, Eudicots, Tracheophytes	Anthocyanidins	Elimination reaction <u>APPLICATION</u> Treatment of larvae of lepidopteran species and influenza virus
7.	Monarda didyma [29]	Lamiaceae Eudicots Tracheophytes	Thymol	Anti seborrheic medicine <u>APPLICATION</u> Treatment for dental caries and gingivitis.
8.	Sambucus Nigra [30]	Adoxaceae Eudicots, Tracheophytes	Beta damascenone Dihydroedulan Phenylacetaldehde Rutinoside	Rutin formation via Quercetin intermediate and phenylalanine starting material <u>APPLICATION:</u> Helpful for the treatment of post-thrombotic syndrome, venous insufficiency, and endothelial dysfunction Treatment Sore throat and conjunctivitis.
9.	Matricaria Chamomilla [31]	Asteraceae Eudicots Tracheophytes	Terpenes Bisabolol Farnesene Chamazulene Flavonoids apigenin Quercetin Patuletin Luteolin Coumarin	Upregulate Gall production in the liver and promotion of emptying of the gall bladder and bile ducts <u>APPLICATION</u> Antiphlogistic Antimutagenic Atropic vaginitis Irritable bowel syndrome

10.	Spilanther acmella [32]	Asteraceae Eudicots Tracheophytes	Spilanthol (2E,6Z,8E)-N-(2 methyl propyl)- 2,6,8 decatrienamide)	Pancreatic lipase inhibitor HIV-1 Protease inhibitor Inhibit sensation of Pain and stimulate saliva secretion due to bitter taste <u>APPLICATION</u> Antinociception Vasorelaxant Throat & gum infection Sialogogue Articular rheumatism
11.	Spilanther Oleracea [33]	Asteraceae Eudicots Tracheophytes	Spilanthol (2E,6Z,8E)-N-(2 methyl propyl)- 2,6,8 decatrienamide)	HIV-1 Protease inhibitor Pancreatic lipase inhibitor <u>APPLICATION</u> Exhibit anti-inflammatory action especially carrageenan-induced paw edema.
12.	Rose damascene mill [34]	Rosaceae Eudicots Tracheophytes	Geranial Citronellol	Manage bacterial population of intestine <u>APPLICATION</u> Antistress activity Relieves uterine congestion causing pain and heavy bleeding.
13.	Achillea millefolium.L [35]	Asteraceae Eudicots Tracheophytes	Linalool Alpha peroxychifolid	Reduce the required blood clotting time and prevent sensory symptoms <u>APPLICATION</u> Skin calming activity
14.	Achillea sibirica ledeb [36], [37]	Lamiaceae Eudicots Tracheophytes	Azulene Caryophyllene Eucalyptol Pinene Borneol Achillene Tannins Caledivain Alkaloids	Stimulate flow of bile & reduce the required blood clotting time <u>APPLICATION</u> Kidney disorders and abscesses, toothaches, amneorrhoea

3. Plant Selection List Based on Oxidate Burst Activity

The following plants are selected based on oxidate burst activity through promoting endogenous immune mediators such as cytokines and chemokines to regulate innate immunity-nonspecific activation in following treatment of immune deficiency conditions including chronic illnesses, sexually transmitted diseases related to liver and spleen, skin disorder color reaction such as leprosy and cell abnormal proliferation such as sarcomas, carcinomas, lymphomas, melanomas

and myelomas through used killed streptococcus pyrogens. In comparative analysis with vaccination especially in cell-mediated humoral immune response producing immunostimulant plants causing nonspecific immune system of larvae of fish and shrimp before specific immune system matures.

Immunostimulant activity will be determined using

- Macrophage migration index (MMI) [38] which is a correlate of macrophage activation and cellular mediated immunity.
- Haemagglutination N or C terminal HA-tagged fusion proteins.
- Plaque forming cell (PFC)count, both of which are the parameter of antigen-specific humoral immunity.
- Derangement of cell-mediated immunity involving T cells in relative to immunological imbalance causing Ig M -Type 4 allergic manifestations and providing evidence of T-Cell alterations in person genetically predisposed to rheumatism. Different types of plants as macrophage migration inhibitory factors are shown in Table 2.

Table 2. Denotes Different Types of Plants ((Macrophage Migration Inhibitory Factor)) Chemical Constituents, Mechanism of Action, and Application.

s.no	LIST OF PLANTS	TAXONOMICAL CLASSIFICATION (Family, Class, Phylum)	CHEMICAL CONSTITUENTS	MECHANISM OF ACTION & APPLICATION
1.	Ocimum sanctum Linn [39]	Lamiaceae Magnoliopsida Tracheophytes	Eugenol Isothymusin Apigenin Circimaritin Cirsilineol Rosameric acid Orientin Vicenin Luteolin Caryophyllene Sesquiterpene Ursolic acid	To balance metabolic stress and chemical stress via normalizing blood glucose concentration, blood pressure, and lipid levels <u>APPLICATION</u> Anxiolytic agent Antidepressant Protect organs and tissues from exposure to cold and excessive noise
2.	Piper longum Linn [40]	Piperaceae Magnolids Tracheophytes	Piperidine Piperonaine Piperoctadecaldine	Reduce platelet aggregation induced by U46619 as a non-competitive thromboxane A2 receptor antagonist <u>APPLICATION</u> Acaricidal activity Fungicidal activity Antiamoebic activity

				Especially helpful for Caecal amoebiasis
3.	Panax Ginseng [41]	Araliaceae Eudicots Tracheophytes	Total ginsenosides Total saponin content in Notoginseng Rhamnogalacturonan I&II, saponin, polysaccharides	Promote neurogenesis by increasing the expression of vascular endothelial growth factor, Brain-derived Neurotropic factor, activating intracellular signaling pathways P13K/Akt and extracellular signal-regulated kinase signaling pathway ERK1/2, inhibiting the hyperphosphorylation of N-methyl D-aspartate receptor 2B. <u>APPLICATION</u> Reduce cerebral ischemia Antivasular aging Anticerebral aging Anticancer properties
4	Withania somnifera [42], [43]	Solanaceae Eudicots Tracheophytes	Isopelletirine anaferine Cuseohygrine, anahygrine, Steroidal lactones withaferin A, withanolides A-Y, Withsomniferin A, Withasomnidienone ,withosmierose A-C, and saponins Sitoindosides and acylsterylglucosides Sitoindosides VII-X and withaferin A (5-dehydroxy withanolide-R)	Exert inhibit effect on ribosome 35S as well as succinate dehydrogenase activity into granulation tissue via stimulation effect on Mg ²⁺ dependant ATPase activity Metabolized component of (someone)withanoside IV helpful to increase neuritis in human neuroblastoma SK-N-Sticells <u>APPLICATION</u> Treatment for Alzheimer's disease Antitumour effect Adaptogenic effect
5.	Mangifera indica linn [44]	Anacardiaceae Eudicots Tracheophytes	Phenolic compounds Xanthones, Flavonols,Gallotannins, Xanthones, Benzophenones, Penta o galloyl-glucose, Ferulic acid Hexoside	To increase absorption and digestion of calcium in the intestine and may hamper in intestinal absorption and assimilation of phosphorus, magnesium,zinc and copper. <u>APPLICATION</u> Antibacterial Antioxidant Antiinflammatory activity

6	Andrographis paniculate burm.f. [45]	Acanthaceae Eudicots Tracheophytes	Diterpenoids Flavonoids Polyphenols Isoandrographolide Neoandrographolide 14 deoxy 11,12 dihydro andrographolide Andrographolide 1	Bind to a spectrum of protein targets including NF-kappa B and actin by covalent modification To increase respiratory infections, fever, Herpes, sore throat and chronic infectious diseases <u>APPLICATION</u> Antiretroviral Antidiabetic Antiangiogenic Antileishmaniasis Antithrombotic Antiurothelithic
7.	Abutilon Indicum linn [46]	Malvaceae Eudicots Tracheophytes	β - Sitosterol	To inhibit the synthesis of tyrosinases and dopaquinone formation <u>APPLICATION</u> Treatment for Gonorrhoea, Septicaemia, leprosy, cystitis and hemorrhagic
8.	Punica granatum Linn [47]	Lythraceae Eudicots Tracheophytes	Punicic acid The conjugated form of α - Linolenic acid	To increase levels of IL-17 IFN- γ in CD 8+ t Cells in mesenteric lymph nodes <u>APPLICATION</u> Chemoprotective activity
9.	Picrorhiza kurroa [48]	Plantaginaceae Eudicots Tracheophytes	Picoside II, III, V 6-Feruloyl catalpol minecoside, Caffeoyl glycoside	To inhibit protein tyrosine kinase activity and stimulate the cell proliferation process of splenocytes, Peritoneal macrophages enhanced cytotoxicity of Natural killer cells. <u>APPLICATION</u> Treatment for Vitiligo, antiperiodic, antiasthmatic, antiperiodic
10.	Phyllanthus Emblica linn [49]	Phyllanthaceae Eudicots Tracheophytes	Kaempferol-3 o- α -L(6 methyl)Rhamnopyranoside(I) Kaempferol-3 o- α -L(6 methyl)Rhamnopyranoside(II)	To reduce expression of Tumour necrosis factor α , interleukin β , intercellular adhesion molecule matrix metalloproteinase -9 inducible nitric oxide synthase, myeloperoxidase. <u>APPLICATION</u> <u>Treatment for</u>

				Leucorrhoea, aphrodisiac, biliousness, asthma and bronchitis
11.	Saussurea costus [50]	Asteraceae Eudicots Tracheophytes	sesquiterpene	To inhibit the mRNA expression of iNOS by lipopolysaccharide-stimulated macrophages, thus reducing nitric acid production <u>APPLICATION</u> Act as cellular signaling, Helps to modulate Vascular tone, insulin secretion, peristalsis, and angiogenesis
12.	Sida cordifolia linn [51]	Malvaceae Eudicots Tracheophytes	Ephedrine, pseudoephedrine, sterculin, β -Sitosterol, Vasicine, saponine, Palmitic acid, Stearic acid	WBC decreased by sister chromatid exchange the exchange of genetic materials between two identical sister chromatids <u>APPLICATION</u> Antiobesity, adaptogenic activity, antigonorrhoea, anti rheumatism as well as treatment for excessive involuntary ejaculation.
13.	Tinospora cordifolia [52]	Menispermaceae Eudicots Tracheophytes	Tinosporide Tinosporine Giloinsterol Sitosterol s, liganin compounds, Giloin Cordifoliside A, B, C & D	To decrease superoxide dismutase, to decrease glutathione peroxidase, To decrease erythrocytes membrane lipid peroxide <u>APPLICATION</u> Anti HIV Antiperiodic activity Anticomplementary activity, antiosteoporotic activity
14.	Wild. T. Sagittata [53], [54]	Lamiaceae Eudicots Tracheophytes	α -Thujone, β caryophyllene, 1,8 cineole, α humulene, β Pinene, Camphor. al lo- aromadendrene, borneol, α -pinene, (z) Salvene & (E) Salvene	Participate in the synthesis of CoQ that is actively involved in electron transport chain oxidative phosphorylation Bas well as dehydrodolichol diphosphate -a precursor of dolichol which transports proteins to the ER lumen for N-glycosylation. <u>APPLICATION</u> To increase growth parameters Act as antimicrobial activity

4. Plant Selection List Based on Management of Autoimmune Disease and Prevention of Transplant Rejections

Immunosuppressants are a structurally and functionally heterogeneous group of drugs that are generally given in combination regimens to treat various autoimmune diseases and to prevent transplant rejections. Immunosuppressants assist the immune system which protects against compartment cell damage and inflammation. The body's immune system helps fight off infections that cause illness. But sometimes immune system mistakenly attacks healthy cells and tissues.

Immunosuppressants can slow down pharmacological responses. This desensitization mechanism was helpful for the treatment of the following diseases such as alopecia areata, inflammatory bowel disease including Crohn's disease and lupus, ulcerative colitis, multiple sclerosis, rheumatoid arthritis, and Psoriasis or psoriatic arthritis) [55]. Immunosuppressants can be recommended for the following diseases inclusive of autoimmune disease, organ transplant, and stem cell or bone marrow transplant [56]. The first choice of treatment for autoimmune disease is Autografts -a kind of medical procedure in which an organ is surgically removed from one body and placed in the body of the recipient to replace a damaged or missing organ. Organ transplantation is used for end-stage organ failures such as liver and heart failure and renal disease. Possibility to do organ transplants composed of kidney, liver, heart, lung, pancreas, intestine, and vascularised composite allograft (VCA) [57] are now possible including face and hand transplantation. Organ transplantation is helpful to patients having disease conditions such as heart disease, diabetes, hepatitis, cystic fibrosis, and cirrhosis. The choice of drug helpful for organic transplants is prednisolone.

The second choice of treatment for autoimmune disease is that stem cell transplants which are used for

- Blood cancers like leukemia, lymphoma, and multiple myeloma.
- Blood disorders like sickle cell disease and thalassemia.
- Bone marrow problems like aplastic anemia.

Allogenic stem cell transplants exchange diseased cells in the human body with healthy ones from a donor called Graft [38]. After transplant donor cells begin to build a new immune system in the human body called the host. The disadvantage of this technique can cause Graft versus host disease (GVHD) to occur in skin, liver, mouth, lungs, Gastrointestinal tract, neuromuscular system or genitourinary tract recipients who have received peripheral blood stem cells/bone marrow from an HLA mismatched related donor have an increased risk of developing acute GVHD. Symptoms for acute GVHD [58]. are skin rash, yellow discoloration of skin and eyes and abnormal blood tests, Nausea, diarrhea or abnormal cramping, and Vomiting.

Symptoms of chronic are rash or discolored areas, skin thickening or tightening abdominal swelling, yellow discoloration of skin and eyes, abnormal blood tests, dry eyes, and vision changes. Dry mouth, white patches inside the mouth, pain or sensitivity to spicy foods, shortness of breath or changes seen on chest X-ray, Difficulty swallowing, pain with swallowing or weight loss, fatigue, vaginal dryness, or pain with intercourse (vagina or vulva) muscle weakness or pain (signs of neuromuscular disease-nerves and muscles), and decreased range of motion in joints or tightness in

joints. prophylactic long-time immunosuppressants are usually used for treatment regimens for chronic graft versus host disease. (GVHD) [59].

Fungal, bacterial, and viral infections are a major risk with this treatment option since your immune system will be desensitized for a long time. Identify the recipient before giving stem cell transplants because it produces potential risk or complications of immunosuppressants.

Blood infections like MRSA and sepsis.

Fungal infections like Thrush and Skin fungus [60].

Skin infections like cellulitis.

Respiratory infections include cold, flu, and pneumonia.

The third choice of treatment for the autoimmune condition is that utilize proper plants, herbs, and nutrients. Different types of plants as immunostimulant activity are shown in Table 3.

Table 3. Denote Different Types of Plants ((Immunostimulant Activity-Rectify and Factor))
Chemical Constituents, Mechanism of Action, and Application.

s.no	LIST OF PLANTS	TAXONOMICAL CLASSIFICATION (Family, Class, Phylum)	CHEMICAL CONSTITUENTS	MECHANISM OF ACTION & APPLICATION
1.	Asparagus racemosus [61]	Asparagaceae Monocots Tracheophytes	Steroidal saponins, Isoflavone, asparagine, racemosol, Polysaccharides, Mucilage, Vitamin A,B1,B2,C,E,Mg, P,Ca,Fe and folic acid Kainic acid	To inhibit oxytocin-induced contractions and stimulate the serotonergic pathway To inhibit Glutathione peroxidase <u>APPLICATION</u> Adaptogenic activity Antihepato Carcinogenesis Antidyspepsia activity Anticancer
2.	Aloe vera [62]	Asphodelaceae Monocots Tracheophytes	Antrokinon Polysaccharide	To decrease the number of WBC and increase phagocytosis in asthmatic adults and to block the reproduction of Herps and AIDS Virus. Alprogen inhibits calcium influx into mast cells thereby inhibiting antigen

				antibody-mediated release of histamine and leukotriene from mast cells. <u>APPLICATION</u> Antivirus, Anti-bacteria, anticancer
3.	Allium sativum [63]	Amaryllidaceae Monocots Tracheophytes	SH Compounds Containing allicin, ajoene, diallyl polysulfides vinyl dithiols, S-allyl cysteine as well as enzymes, saponins and flavonoids	It promotes caspase 3 activity and increases ROS in cancer cells and causes a reduction in Myocardial ischemia-reperfusion injury and increases eNOS (Endothelial nitric oxide synthase) activation via phosphorylation by H ₂ S through activation of the P13K/AKT pathway and promotes mitochondria function negatively. <u>APPLICATION</u> Treatment for prostate and breast cancer.
4.	Azadirachta indica A.juss [64], [65]	Meliaceae Eudicots Tracheophytes	Flavonoids, alkaloids, tannins, saponins, and steroids	Reduction reaction happened on urea concentration <u>APPLICATION</u> Antioxidant, Anticancer and anti-inflammatory
5.	Andrographis paniculata burn F [66]	Acanthaceae Eudicots Tracheophytes	Diterpenoids Flavonoids Polyphenols Isoandrographolide Neoandrographolide 14 deoxy 11,12 dihydro andrographolide Andrographolide 1	Bind to a spectrum of protein targets including NF-kappa B and actin by covalent modification To increase respiratory infections, fever, Herpes, sore throat, and chronic infectious diseases <u>APPLICATION</u> Antiretroviral Antidiabetic Antiangiogenic

				Antileishmaniasis Antithrombotic Antiurothelithic
6.	Albution indicum lin [67]	Malvaceae Eudicots Tracheophytes	β - Sitosterol	To inhibit the synthesis of tyrosinases and dopaquinone formation <u>APPLICATION</u> Treatment for Gonorrhoea, Septicaemia, leprosy, cystitis and hemorrhagic
7.	Cynodon dactylon linn [68]	Poaceae Monocots Tracheophytes	Crude protein carbohydrate	To decrease the number of WBC and increase phagocytosis in adults and to block the reproduction of Herps and AIDS Virus. It involves in nitric oxide scavenging action <u>APPLICATION</u> Treatment for Aluminium induced neurotoxicity Carbofuran-induced oxidative stress and white spot syndrome
8.	Curcuma long [69]	Zingiberaceae Monocots Tracheophytes	α - Tumerone C3 -curcuminoids C2- Tumerone	Exhibit stimulatory effects on PBMC proliferation and cytokine production. <u>APPLICATION</u> Anticancer activity Depurative Carminative Treatment for sprains and bruises. Antiplatelet activity Antiproliferative effect
9.	Embelia ribs burm [70]	Primulaceae Eudicots Tracheophytes	Stigmasterol Sitosterol Embelin Taraxasterol Betulin	It decreases paraquat-induced lung injury by suppressing oxidative stress, inflammatory cascade (inflammatory cytokine release), and MAPK/NF- κ B Signaling pathway in

				paraquat intoxicated animals. Embelin and embelin derivatives selectively inhibit 5-LOX and microsomal prostaglandin E2 synthase I. <u>APPLICATION</u> Anthelmintic Astringent Carminative Antifertility Treatment for Ascariasis
10.	Nuctanther arbor tristis Linn [71]	Oleaceae Eudicots Tracheophytes	Oleanolic acid (Betulinic acid) Glycoside Naringenin Iridoid glycoside Arbortistosides A,B &C	C3 -Esterification of betulinic acid acts as HIV 1 maturation inhibition via the mevalonate pathway <u>APPLICATION</u> Antisnake venom activity Treatment for sciatica, arthritis, fever and laxative Anticancer activity

5. Plant Selection List Based on Blocking Macrophage Inhibitory Factor Activity

MIF gene is located in chromosome 22q 11.2 and is a highly conserved 12.5 KDa protein that implies a unique combination of hormone-like, cytokine, and thioredoxin-like properties and is recognized to be a multipotent cytokine involved in the regulation of immune and inflammatory responses via various mediators and plays a key role in regulating several inflammatory and immune-mediated diseases seen in pannus, kidney, gut and brain and blood vessels. The identification of single nucleotide polymorphism at position 173 (MIF 173C allele) and CATT5-8 tetranucleotide repeat element of the MIF gene has sparked research [72].

Empathetic pathogenesis factor leading to the cause of C-reactive protein in the unbalanced situation can produce rheumatic diseases such as rheumatoid arthritis, systemic vasculitis, and systemic lupus erythematosus in association with an autoimmune disease in coupling with rheumatic diseases.

5.1 Interpret evaluation for attaining amplification of pro-inflammatory loop

Maintain Glucocorticoids at optimum levels either increased or decreased concentration of Glucocorticoids will tend to alter cellular ATP dependant concentration and cause infectious disease, storage disease, congenital disorder, and chromosome disorder.

- Upon upregulation stimulation reaction cytokines, LPS, TLRs are stimulated which are located in sites pannus, blood vessels, kidney, gut, and brain- corresponding release of Glucocorticoids from Hypothalamus -pituitary-adrenal axis through playing MIF in regulating global responses to infection.
- Hyper increment of CRP will lead to increase synovitis due to inflammatory pannus, the proliferation of synoviocytes, infiltration by inflammation and immune cells as well as increased MIF levels in synovial fluid and synovial tissue especially in humans having Juvenile idiopathic arthritis.
- Hypo increment of CRP will lead to cause myocarditis.
- C-Reactive protein will be maintained at optimum level in the following aspects likewise no heart failure, no invasion of foreign or microorganism entry, no pathogenic activity, and no complications.
- Neutralisation mechanism which enhances protective effects against lethal bacterial sepsis and septic shock induced by Gr negative endotoxin and Gr positive exotoxin, especially in MIF deficiency.

Hyper increment of CRP in patients can be tested by psychiatric problems with emotional stress, normal behavioral changes, inability to do sex, loss of libido or ability to think to do sex but condition impossible, and rigidity of morning erection. This can be altered through exercise, yoga, meditation, and holding high weight.

Rectify MIF block activity in sepsis condition either through

- MIF gene alteration
- Anti-MIF antibodies production
- Decrease cytokine production by down-regulating Toll-like receptor 4 expression
- Pro-inflammatory molecules TNF- α , IL-5, IFN- γ transforming growth factor β , and liposaccharide (LPS) have been shown to stimulate MIF mRNA expression and protein secretion.
- Complement activated protein C5a promotes MIF release from PMN in vitro and during sepsis.
- Remedy to prevent rheumatic arthritis [73] with sepsis [74] condition patients, increase binding capacity with TLR4 ligands to increase MIF production.
- Remedy to prevent rheumatic arthritis with HSV-2 Herpes simplex virus condition having focal necrotizing hepatitis condition to activate macrophages and dendritic cells in liver, spleen, skin and lungs [75].

6. Research Hypothesis Regarding Antibiotic Resistance in Relation to Auto-Immune Diseases or Disorders

Frequent antibiotics such as Clindamycin, Trimethoprim, Amoxicillin, and Ciprofloxacin are used in treating lymphadenitis, lympharngitis, and scrofula diseases. Treatment will be affected by antibacterial medications. But these antibiotics cause resistance in correlation with Idiopathic and self-limiting ADR which causes toxicity such as bacterial or viral infections, Tuberculosis lymphadenitis, and cat-scratch diseases. These identification factors will tend to recognize that the person is affected by a viral infection, not for bacterial infection. These superinfections due to antibiotic resistance tend to cause the following symptoms as like as Quincke's edema [76] (urticaria, redness edema), change wounds, runny nose, inflammation of the conjunctiva, fever, joint pain, eosinophilia serious complications, Anaphylactic shock. Fungal lesions of the oral cavity, intestinal dysbiosis, vagina, digestive disorders, Violations from the nervous system, increased reflex excitability, signs of meningism, convulsive conditions, coma, Neurological

manifestations, violation of blood clotting, water-electrolyte disorders in the form of hypokalemia, hyponatremia. Disorders of the urinary system such as the appearance of hematuria, proteinuria, the development of intestinal nephritis, burning and redness of the skin and cholestatic jaundice [77], [78].

6.1 The formation of superinfection in systemic treatment can be prevented by following selected plants through mechanism

6.1.1 Our lymphatic systems are responsible for removing cellular waste in our bodies.

- a) Everything from mutated cells to viruses and bacteria are removed by the lymphatic system.
- b) when lymph becomes sluggish and congested, human bodies have a hard time keeping up with all of the waste, environmental toxins, viruses etc that happens in contact relationships who have not have optimal health.
- c) Lymph's sluggish human damage system is recognized by following signs and symptoms such as fatigue, breast swelling with each cycle, itchy and dry skin, bloating, PMS symptoms in women, stress, exhaustion, clouding of consciousness-Brain fog, [79], [80] swollen glands, stubborn weight gain, Chronic sinusitis, sore throat, colds or ear issues, Chronic serious congestion, Cellulite, Cold hands and feet's.
- a) Herbs that help to promote lymphatic drainage activity:
- b) Activity herbs selected to Rectify Antibiotic Resistance-ADR.

Different types of plants as the detoxifying lymphatic system are shown in Table 4.

Table 4. Denotes Different Types of Plants ((Detoxifying Lymphatic System) Chemical Constituents, Mechanism of Action, and Application.

s.no	LIST OF PLANTS	TAXONOMICAL CLASSIFICATION (Family, Class, Phylum)	CHEMICAL CONSTITUENTS	MECHANISM OF ACTION & APPLICATION
1.	Gallium aparine [81]	Rubiaceae Eudicots Tracheophytes	Iridoid glycosides Coumarin alkaloids Anthraquinone aldehyde	<u>MOA:</u> NADPH dependant reduction step followed by cyclization step that occurs through either Diels alder reaction or intramolecular Michael addition <u>Application:</u> Antifeedant activity against spodoptera litura
2.	Calendula officinalis [82]	Asteraceae Eudicots Tracheophytes	Flavonoid glycosides Triterpene Oligoglycosides Oleanna type Triterpene glycosides Saponins Sesquiterpene Glycosides	<u>MOA</u> To inhibit HIV-1 Replication of encephalitis virus, herpes simplex virus, influenza APR-8 virus. <u>Application:</u> Antimutagenic

				To stimulate cellular metabolism by an increase of mitochondrial dehydrogenase activity.
3.	Echinacea angustifolia [83]	Asteraceae Eudicots Tracheophytes	Cis and Trans Traumatic acid salts Cichoric acid Cynarine	<u>MOA</u> To promote wound healing polysaccharide fraction (Ehinacin B)healing by forming a hyaluronic acid polysaccharide complex that indirectly leads to inhibition of hyaluronidase and stimulates the growth of fibroblasts. <u>Application:</u> cicatrizant activity Antihyaluronidases, To inhibit arachidonic acid metabolism to inflammatory prostaglandins
4.	Astragalus membranaceus [84]	Fabaceae Eudicots Tracheophytes	Saponins Isoflavone flavonoids	<u>MOA</u> To increase Telomerase activity To inhibit oxidant stress by upregulating antioxidant factors To upregulate expression of nuclear factor erythroid 2 related factor, SOD, catalase, glutathione peroxidase, To decrease reactive oxygen species, To decrease the production of malondialdehyde and free radical levels, and reduce cell apoptosis. <u>APPLICATION</u> Antivascular aging, Anti-brain aging, Anti-inflammatory, Anti-cancer effects Hepatoprotective
5.	Taraxacum officinalis [85]	Asteraceae Eudicots Tracheophytes	Saikosaponins Triterpene saponin glycosides	<u>MOA</u> To attenuate proinflammatory cytokine production

				To inhibit viral replication through downregulating NF-KB signaling caspase 3 dependant virus ribonucleoprotein nuclear export, neutrophil and monocytes recruitment. <u>APPLICATION:</u> Hepatoprotective immunomodulators reduce viral load as well as inhibitory viral replication activity.
6.	Scrophularia nodosa [86]	Scrophulariaceae Eudicots Tracheophytes	Saponins Tannins Phytolectin	<u>MOA</u> Cell differentiation and urease inhibition <u>APPLICATION</u> Tubercular testis and protruding piles Breast tumors
7.	Lycopodium clavatum [87]	Lycopodiaceae Lycophtes Tracheophytes	Alkaloids Triterpenoids	<u>MOA</u> To inhibit acetyl cholinesterase activity, <u>APPLICATION</u> Antioxidative Antiproliferative activity for HepG2 Cells
8.	Harpagophytum procumbens [88]	Pedaliaceae Eudicots Tracheophytes	Harpagoside Iridoid glycoside Phytosterols flavonoids	<u>MOA</u> It inhibits inflammatory mediators including cyclooxygenase 2, leukotrienes, nitric oxide, Tumour necrosis factor - α , and interleukin - β and by inhibiting metalloproteinases and elastase. <u>APPLICATION</u> Treatment for AIDS-induced fever, allergy.
9.	Hydrastis Canadensis [89,90]	Ranunculaceae Eudicots Tracheophytes	Isoquinoline alkaloids Berberine Berberastine Tetrahydroberberastine Hydrastine Hydrastinine Canadine canalidine	<u>MOA</u> It inhibits the activity of Topoisomerase I & II It inhibits CYP3A4, CYP2C9,CYP2D6,CYP2C19 <u>APPLICATION</u> Antiarrhythmics To improve blood circulation in defects such as

				piles, varicose veins, and edema. Treatment for Hay fever and upper respiratory problems.
10.	Baptisia tinctoria [91]	Fabaceae Eudicots Tracheophytes	Alkaloids Steroids Flavonoids Triterpenoids Coumarins Tannins	It helps to inhibit the parasympathetic nervous system b selectively blocking the binding of ACH synapses in CNS & PNS to its receptor in nerve cells. <u>APPLICATION</u> Anticonvulsant activity Antistress activity Antianxiety activity
11.	Mullein Verbascum Thapsus [92]	Scrophulariaceae Eudicots Tracheophytes	Saponins Iridoid glycosides Phenylethanoid glycosides Flavonoids Vitamin c Minerals	<u>MOA</u> Exhibit antihepatoma activity on five human liver cancer cell lines HepG2/C3A, SK-HEP-1, HA22T/VGH, Hep3b & PLC/PRF/5 <u>APPLICATION</u> Treatment for Rey's syndrome and inner ear infection.
12.	Glycyrrhiza glabra [93]	Fabaceae Eudicots Tracheophytes	Triterpenoid saponin glycoside	<u>MOA</u> Decrease serum testosterone level in women and beneficial in aplastic anemia <u>APPLICATION</u> Detoxify and protects the liver. Treatment for Leucorrhoea and Addisons diseases
13.	Trifolium pretense.L [94]	Fabaceae Eudicots Tracheophytes	Isoflavones Formonnectin Biochanin A Daidzin Genistein	<u>MOA</u> It inhibits aromatase activity and expression To purify the blood promoting urine, mucous and bile production and improving circulation. <u>APPLICATION</u> Menopause treatment to balance hot flashes Anti-inflammatory Antioestrogenic activity

14.	Commiphora wightii [43]	Burseraceae Eudicots Tracheo-phytes	Guggulsterone Quinic acid Myo inositol	<u>MOA</u> Acts as an antagonist of the Farnesoid X receptor and reduces cholesterol synthesis in liver <u>APPLICATION</u> It mediates cell signal transduction in response variety of hormones, neurotransmitters and growth factors and participates in osmoregulation
15.	Ashwagandha [43]	Solanaceae Eudicots Tracheophytes	Isopelletirine anaferine Cuseohygrine, anahygrine, Steroidal lactones withaferins and saponins Sitoindosides and acylsterylglucosides Sitoindosides VII-X and withaferin A (5- dehydroxy withanolide- R)	Exert inhibit effect on ribosome 35S as well as succinate dehydrogenase activity into granulation tissue via stimulation effect on Mg 2+ dependant ATPase activity Metabolized component of (sominone)withanoside IV helpful to increase neuritis in human neuroblastoma SK-N-Sticells <u>APPLICATION</u> Treatment for Alzheimers disease Antitumour effect Adaptogenic effect
16.	Curcumin plus kanchanaar guggul [43]	Zingiberaceae Monocots Tracheophytes	α - Tumerone	Exhibit stimulatory effects on PBMC proliferation and cytokine production. <u>APPLICATION</u> It intervenes in dispute cell signal transduction in response variety of hormones, neurotransmitters and growth factors.
17.	Heliotropium Indicum [96]	Boraginaceae Eudicots Tracheo-phytes	Indicine principal base, Echinatine Supinine Heliurine Helitrine Lasiocarpine Lasiocarpine	<u>MOA</u> Formation of microsomal oxidation of Lasiocarpine (dehydroheliotridine) Undergoes replacement reaction to remove poisonous materials such as

			N-oxide	snake bites and scorpion stings. <u>APPLICATION</u> Antidote Helpful for treatment of Thrush, skin ulcers and boil furuncle, and Ocular infections
18.	Rhaphoticum carthamoides [97]	Asteraceae Eudicots Tracheophytes	Ecdysteroids Flavonoids Phenolic acids	<u>MOA</u> Acts as a free-radical scavenger and reduces the amount of damage from ROS It reduces Glutamate induced excitotoxicity in the hippocampus. <u>APPLICATION</u> Antibraincancer Antibreastcancer Anticervicalcancer Helps endurance strength and recovery

6. Conclusion

Hidradenitis suppurative is a kind of chronic inflammatory disease in association with modifying co-morbidities including metabolic syndrome, androgen dysfunction, and smoking. In addition to correlates with psychological factors that are linked with various kinds of diseases such as apocrine gland dysfunction, follicular occlusive diseases, comorbidities linked with autoimmune diseases and proliferative cells, depression, hyperlipidemia, acne, obesity, Insulin resistance, pilonidal sinus, polycystic ovary syndrome, diabetes, Hypertension and keratosis pilaris. Among Ig various types, Ig A plays a key role in different inflammatory conditions and auto-immune conditions. Ig A produces kickball change, especially in mucosal hemostasis such as GI, respiratory and Genitourinary tracts. Impairment O – linked glycosylation process during glomerular injury tends to produce an immune complex which is prevented by Gym practice exercise using the instrument. A sexually transmitted disease-oriented neurological complication is managed with Tai chi stagnant and Tai chi Qigong motion movement.

EVPM cannot suitable for JEB (Junctional epidermolysis bullosa) disease conditions. But these conditions are managed with well-balanced diet and Yoga meditations. Cognitive-based therapy is helpful for the management of eight intellectual domains such as memory, language, perceptual skills, attention, motor skills, orientation, problem-solving skills, and executive functioning abilities.

Comparative analysis of evidence-based plant medicine with the allopathic drug in viral infections is evaluated based on different factorial designs of lymphadenopathy and various conditions. The reason behind EBPM is the right choice

comparatively allopathic medicine which is producing various side effects such as bronchospasm, stuffed nose, running nose, behavioral changes, diarrhea, cough, nausea, vomiting, cold sores, shaky movements due to oscillatory muscle contraction-unintentional rhythmic movements, hallucinations, coma, confusion, less urine, severe drowsiness, dry mouth, muscle pain, and skin rash.

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