# Pharmaceutical standardization, product development and quality control aspect of C- Pills a marketed

# **Product**

A REPORT

SUBMITTED IN PARTIAL FULFILLMENT OF THE REQUIREMENTS FOR THE DEGREE OF

## MASTER OF PHARMACY (AYURVEDA)

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#### Introduction

In the present era, the herbal drugs have gained fame all over the world. Indian medicine system is one of the oldest medicine systems in the world and it has the number of herbs and herbal formulation to treat the various disorders. Dengue is recognized to be the most significant arthropodborne viral disease of humans worldwide  $^{(1)}$ . Number of peoples is killed by the every year and there is no effective medicine is available for the dengue. The disease has also been termed "break bone fever" or "dandy fever" and in Ayurveda as *dandak jwara*. In Ayurveda, jwara is very complex diseases there are 80 types of fevers are mention. Explanation of this is found in "*madav nidan*" under the chapter *parishishta* chapter.  $^{(2)}$  Dengue fever begins with a sudden high fever, often as high as 104 - 105 degrees, 4 to 7 days after the infection. Headache, retro orbital pain, and back pain along with the severe Muscular pain that gave rise to the colloquial designation "break-bone fever." A flat, red rash may appear over most of the body 2 - 5 days after the fever starts.  $^{(3)}$ 

C-pill is a combination of herbs which can help to increase the platelet count during the dengue or other platelets related complications. It has hepatoprotective activity and also protects the vital system of the body. It also gives strength to immune system by enhancing both cellular and humoral immunity. It also prevents the platelets aggregation and help to control the fever.

This is contraindicated during the pregnancy, it include the *carica papaya* as ingredient which may have deal to the miscarriage. C-pill has no side effect which is reported. These are remanded to take minimum 10 days for good result.

#### **SCOPE OF STUDY**

C-pills are the marketed product which has shown effective response against complications associated with dengue. Herbal combination shows a significant response against the DENV1 type and help to increase the platelet counts. It also showed the immunomodulator activity due to its antioxidant properties. The combination of herbal material not shown a significant efficacy against other types of dengue virus i.e. DENV2, DENV3, DENV4, DENV5; So there is a need to improve formula which may lead the effective treatment against dengue virus. The study is aimed to improve the formula provide the effective response against all type dengue complication and standrization of prepared formulation (tablet, capsule, syrup, phytosome). These are some various issues which are of concern

#### **OBJECTIVE OF STUDY**

- To perform standard parameters for the standardization of the raw material
- Development of various dosage form which can be beneficial for the dengue patients
- Pharmaceutical standardization and stability study of prepared dosage form
- To carry out the comparative study of prepared dosage form

#### LITERATURE REVIEW

#### 4.1 Review of the dengue

#### 4.1.1 According to Ayurveda

In Ayurveda fever is called as Jwara. Also have different synonyms like Kshaya because it destroys body as well as Dhatu. Jwara is also called as Tama because it produces moha which is the function of tama. Jwara is also called as Yamatmaka because it kills the person like that of Yamaraja. There are number of jwara are mention dengue is one of them and known as "dandak jwara". Dengue is also called as hadjod jwara and dandy fever. (3) (4)

#### **Causative factor (Nidan)**

Main cause of Dandak jwara is Dengue virus which spread in body through Aedes Aegypti mosquito.

#### **Premonitory symptoms (Purvarupa)**

Angmard- bodyache, klam -tiredness without exertion, aruchi- anorexia, nausea, avsaad-depression.

#### Symptoms (Rupa)

Severe breaking pain in bone and joints. High temperature of 103 to 105 degrees F. may occur which gets subside and may relapse again within three to four days (Saddle back fever). On 8th day, it subsides on its own. Severe pains in bones, difficulty in walking, slow pulse, excessive weakness, loss of appetite are common symptoms. During fever, pulse is not proportionately as fast as it should be with fever. Symptoms of common cold (Pratishyaya)

cough and throat pain are also common symptoms of Dandaka jwara which becomes endemic due to virulence of kapha and vata dosha. (5)

#### **Etiology (Hetu)**

According to Ayurveda, disease is caused only when there is lack of immunity in the body. In this ancient science of healing, when the human body contains ama, it becomes a fertile breeding ground for different kinds of viruses. In the human body ojas is the product of a healthy digestive system, which helps the body in resisting infection. Low ojas is lack of immunity at the physical level and absence of mental strength at the mind level. Dengue can affect anyone but tends to be more severe in people with compromised immune systems.

Due to Virus infestation and irregular nature of fever it can be co-related with dandak Jwara described in Ayurvedic text. Hetu of dandak Jwara has been described. Intermittent fever as caused by invasion of evil spirits or micro-organisms. <sup>(6)</sup>

#### Pathogenesis (Samprapti)

A condition in which the body temperature goes beyond the normal temperature and is characterized by disturbance in normal functioning of the system. Fever occurs when the digestive fire (agni) and digestive toxins (ama) which are normally found in the gastrointestinal tract are change their place by disrupted doshas and then they overflow into the blood and lymphatic system.<sup>(6)</sup>

#### Management

Ayurveda many medicinal plants are effective in Dengue fever and such a very impotant role in cure to Dengue fever. Ayurvedic medicines, herbs, rasayanas and much Ayurvedic therapy are use in treat for dengue fever.

According to Ayurveda spread of dengue can be prevented by strengthening the ojas and enhancing body immunity. Ayurveda is a practical, medical science which promotes perfect health through prevention and cure health problems by recommending lifestyle changes. Ayurveda, hygiene, lifestyle and diet are crucial to good health. To attain good health, Ayurveda prescribes the specific daily routine Dincharya and the seasonal regime Ritucharya. These are the some drugs which can be used for the treatment of the dengue. (3)(6)

#### **4.1.2** According to the Modern science

Dengue virus (DENV) is the cause of dengue fever. It is a mosquito-borne single positive-stranded RNA virus of the family Flaviviridae; genus Flavivirus. Five serotypes of the virus have been found, all of which can cause the full spectrum of disease. Dengue fever is a mosquito-borne tropical disease caused by the dengue virus. Symptoms typically begin three to fourteen days after infection. This may include a high fever, headache, vomiting, muscle and joint pains, and a characteristic skin rash. Recovery generally takes two to seven days. In a small proportion of cases, the disease develops into the life-threatening. Dengue hemorrhagic fever, resulting in bleeding, low levels of blood platelets and blood plasma leakage, or into dengue shock syndrome, where dangerously low blood pressure occurs.<sup>(7)</sup>

**Virology:** Dengue virus is a RNA virus of the family flaviviridae3; they are otherwise called arboviruses. The dengue virus genome contains 11,000 nucleotide bones. They have 3 different

protein molecules that form virus partied (C, prM and E) and 7 other types of protein molecules (NSI, NS2a, NS2b, NS3, NS4a, NS4b, NS5) that are found in infected host cells and are required for replication of virus. There are 4 strains of virus, ex; DEN1, DEN2, DEN3, DEN4. ALL 4 serotypes can cause full blown disease. Infection with 1 serotype is believed to produce lifelong immunity to that serotype, but he can be infected with other serotypes in future. (8)(9)(11)

The humans are the primary host for dengue viruses & transmitted by Aedes mosquitoes. A mosquito that takes a blood meal from infected person become infected with virus. In 8 to 10 days.

#### Mechanism

When a mosquito carrying DENV bites a person, the virus enters the skin together with the mosquito's saliva. It binds to and enters the white blood cells, and reproduces inside the cells while they move throughout the body. The white blood cells respond by producing a number of signal proteins (Such as interferon) that are responsible for many of the symptoms, such as the fever, the flu-like symptoms and the severe pains. In severe infection, the virus production inside the body is much increased, and many more organs (Such as the liver and the bone marrow) can be affected, and fluid from the bloodstream leaks through the wall of small blood vessels into body cavities. As a result, less blood circulates in the blood vessels, and the blood pressure becomes so low that it cannot supply sufficient blood to vital organs. Furthermore, dysfunction of the bone marrow leads to reduced numbers of platelets, which are necessary for effective blood clotting; this increases the risk of bleeding. (10)(11)

#### **Clinical Features**

The characteristic symptoms of dengue are: a sudden-onset fever, headache (Typically behind the eyes), muscle and joint pains, and a rash; the alternative name for dengue, "break-bone fever", comes from the associated muscle and joints pains. The course of infection is divided into three phases: febrile, critical, and recovery.

The febrile phase involves high fevers, frequently over 40°C (104°F) and associated with generalized pain and a headache; this usually lasts 2–7 days. Flushed skin and some small red spots called petechiae, which are caused by broken capillaries, may occur at this point.

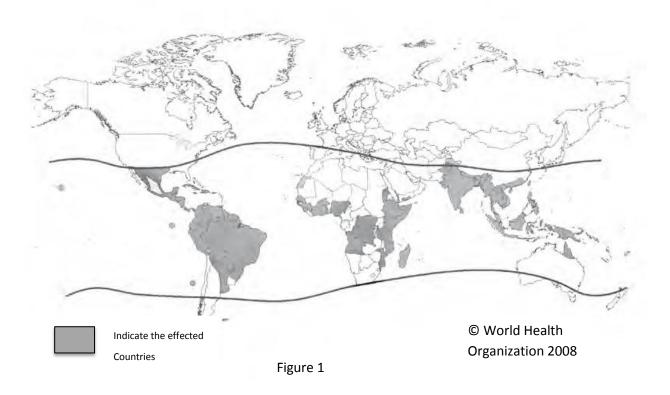
The critical phase, if it occurs, follows the resolution of the high fever and typically lasts one to two days. During this phase there may be significant fluid accumulation in the chest and abdominal cavity due to increased capillary permeability and leakage. This leads to depletion of

fluid from the circulation and organs. During this phase, organ dysfunction and severe bleeding (Typically from the gastrointestinal tract) may occur. Shock and hemorrhage occur in less than 5% of all cases of dengue but those who have previously been infected with other serotypes of dengue virus ("Secondary infection") have an increased risk of this.

The recovery phase occurs next, with resorption of the leaked fluid into the bloodstream. This usually occurs over a period of two to three days. The improvement is often striking, but there may be severe itching and a rate. It is during this stage that a fluid overload state may occur, which if it affects the brain may reduce the level of consciousness or cause seizures. (11)(19)

#### 4.1.3 PROBLEM BACKGROUND

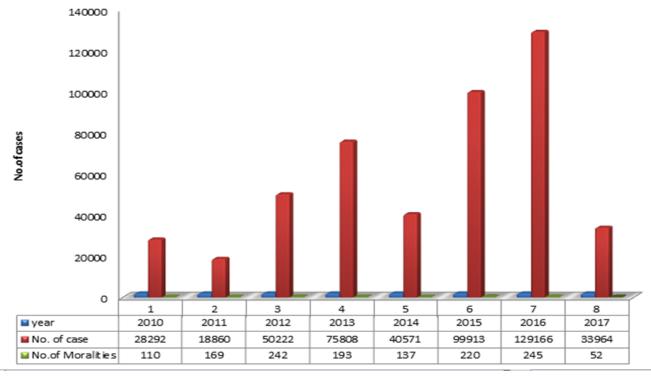
Dengue is the most rapidly spreading mosquito-borne viral disease in the world. In the last 50 years, incidence has increased 30-fold with increasing geographic expansion to new countries and, in the present decade, from urban to rural settings. An estimated 50 million dengue infections occur annually and approximately 2.5 billion people live in dengue endemic countries.



#### **Dengue** in india

Dengue is the most rapidly spreading mosquito-borne viral disease in the world. India reported 1.3 million cases in 2006 but no data on mortality were available. The history of dengue outbreaks in India has been recently reviewed. More recent and systematic data are now available because of the NVBDCP. The data on the web site of NVBDCP.

According to National Vector Borne Disease Control Program data from 2010 there are number of cases decrease or increase but the number mortality is decrease against the number of cases reported per year till 2017. That is possible due to the more awareness of people to dengue or the improvement in the medication of this disease . but we have to aware peoples more about this disease and about the preventions of it . so we can decrease the mortality in the coming years. (12)



Reported cases and mortalities from 2010 to 2017  $^{(13)}$ 

#### Contraindication

Dengue presents as a fever with body ache in the beginning, most people mistake it as a case of viral fever. They end up self-medicating themselves with either Aspirin or Ibuprofen. Now, this can be very dangerous and cause severe bleeding. (14)

#### 4.2 Review of the product

Shree Dhanwantri Herbals was founded by Vaidya Deen Dayal Singh Ji in the year 1952. The Pharmacy which started in a very humble way has grown by leaps and bounds under the able guidance of its directors, and tireless and sincere efforts put in by the workers in these years. C-Pill Tablets are manufactured by Shree Dhanwantri Herbals. Shree Dhanwantri C-Pill Tablets has composition of Eranda Karkati and Tulsi. C-Pill Tablets supports for fever.

#### 4.3 Raw material review

#### 4.3.1 Tulsi (Ocmium sanctum)

#### • API (The Ayurvedic Pharmacopeia of India)

Tulsi consists of dried whole plant of <u>Ocimum sanctum Linn</u>. (Fam. Lamiaceae) An erect, 30 - 60 cm high, much branched, annual herb, found throughout the country. (15)(16)

#### **Synonyms**

Language	Synonynms					
Sanskrit	Surasa, atulase, Bana Tulasi					
Assamese	Tulasi					
Bengali	Tulasi					
English	Holy Basil					
Gujrati	Tulasi, Tulsi					
Hindi	Tulasi					
Kannada	Tulasi, Shree Tulasi, Vishnu					
Tulasi						
Malayalam	Tulasi, Tulasa					
Marathi	Tulas					

Punjabi	Tulasi
Tamil	Tulasi, Thulasi, ThiruTheezai
Telugu	Tulasi
Urdu	Raihan, Tulsi

#### **Description**

#### a) Macroscopic

#### Root

Thin, wiry, branched, hairy, soft, blackish-brown externally and pale. Violet Internally.

Stem - Erect, herbaceous, woody, branched hairy, sub quadrangular, externally Purplish-brown to black, internally cream, coloured, fracture, fibrous in bark and short in Xylem, dour faintly aromatic.

#### Leaf

2.5-5 cm long 1.6 - 3.2 cm wide, elliptic oblong, obtuse or acute, entire or serrate, Pubescent on both sides; petiole thin, about 1.5-3 cm long hairy, odour, aromatic taste Characteristic.

#### **Flower**

Purplish or crimson coloured, small in close whorls; bracts about 3 mm longand broad, pedicels longer than calyx, slender, pubescent; calyx ovoid or campanulate3-4 mm bilipped, upper lip broadly obovate or suborbicular, shortly apiculate, lower lip longer than upper having four mucronate teeth, lateral two short and central two largest, corolla about 4 mm long, pubescent; odour, aromatic; taste, pungent.

#### Fruit

A group of 4 nutlets, each with one seed, enclosed in an enlarged, membranous, veined calyx, nutlets sub-globose or broadly elliptic, slightly compressed, nearly smooth; pale brown or reddish with small black marking at the place of attachment to the thalamus; odour, aromatic; taste, pungent.

#### Seed

Rounded to oval; brown, mucilaginous when soaked in water, 0.1 cm long, slightly notched at the base; no odour; taste, pungent, slightly mucilaginous.

#### b) Microscopy

#### Root

Shows a single layered epidermis followed by cortex, consisting of seven or more layers of rectangular, round to oval polygonal, thin-walled, parenchymatous cells, filled with brown content, inner layers of cortex devoid of contents; phloem consisting of sieve elements, thin-walled, rectangular parenchyma cells and scattered groups of fibres, found scattered in phloem; xylem consists of vessels, tracheids, fibres and parenchyma, vessels pitted; fibre tracheides, long, pitted with pointed ends; fibres thick walled and with pointed ends.

#### Stem

Shows a single layered epidermis with uniseriate, multicellular covering trichomes having 5-6 cells, occasionally a few cells collapsed; cortex consists of 10 or more layers of thin-walled, rectangular, parenchymatous cells; phloem consists of sieve elements, thin-walled, rectangular parenchyma cells and fibres; fibres found scattered mostly throughout phloem, in groups and rarely in singles; xylem occupies major portion of stem consisting of vessels, tracheids fibres and parenchyma; vessels pitted, fibres with pointed ends; centr; pith consisting of round to oval, thin-walled, parenchymatous cells.

#### **Leaf-Petiole**

Shows somewhat cordate outline, consisting of single layered epidermis composed of thin-walled, oval cells having a number of covering and glandular trichomes; covering trichomes multicellular 1-8 celled long, rarely slightly reflexed at tip, glandular trichomes short, sessile with 1-2 celled stalk and 2-8 celled balloon shaped head, measuring 22-27 in dia; epidermis followed by 1 or 2 layers and 2 or 3 layers of thin-walled, elongated, parenchyma cells towards upper and lower surface srespectively; three vascular bundles situated centrally, middle one larger than other two, xylem surrounded by phloem.

#### Midrib

Epidermis, trichomes and vascular bundles similar to those of petiole except cortical layers reduced towards apical region. Lamina - epidermis and trichomes similar to those of petiole; both anomocytic and diacytic type of stomata present on both surfaces, slightly raised above the level of epidermis; palisade single layered followed by 4-6 layers of closely packed spongy

parenchyma with chloroplast and oleo-resin; stomatal index 10-12-15 on upper surface and 14 - 15 - 16 on lower surface; palisade ratio 3.8; vein islet number 31 - 35.

#### **Powder**

Greenish: shows thin-walled, parenchymatous cells, a few containing reddish brown contents, unicellular and Multi-cellular trichomes either entire or in pieces; thin walled fibres, xylem vessels with pitted thickenings, fragments of epidermal cells insurface view having irregular shape, oil globules, rounded to oval, simple as well as compound starch grains having 2-5 components, measuring  $3-17~\mu$  in diameter.

#### **IDENTITY, PURITY AND STRENGTH**

Foreign matter

Not more than 2 per cent.

Not more than 10 per cent.

Not more than 1.5 per cent.

Not more than 1.5 per cent.

Not less than 4 per cent.

Water-soluble extractive

Not less than 8 per cent.

#### T.L.C.

T.L.C. of Tulasi oil obtained by stem distillation is carried out on Silica gel 'G' plate using Toluene :Ethyl acetate (93:7) Tulasi oil is diluted in chloroform-toluene (1:10). Eugenol to be applied as standard also diluted in 130 ratio and 10 µl of each to beapplied in band form. After running distance of 10 cm the plate is air drying for 15minutes and then kept in the oven for 2 to 5 minutes. On cooling spray, in thoroughly vanillin- Sulphuric acid reagent and heat the plate at 110° C for 5 - 1- minutes under observation. Record Rf. values of eugenol and caryophyllence. Eugenol (orange brown) approx. Rf. value 0.7, caryophyllence (reddish violet) runs to solvent front.

#### **Constituents**

Essential Oil.

#### **Properties and action**

Characters	Properties			
Rasa	Katu, Tikta, Kashya			
Guna	Laghu, Ruksha			
Virya	Ushna			

Vipaka	Katu					
Karma	Depana, Hirdya, Kaphahara, Rucya,					
	Vatahara, Pittavardhine, Durgandhihara					

#### **Important formulations**

Tribhuvanakerti Rasa, Muktapanchamrta Rasa, MuktadiMahaanjana

#### Therapeutic uses

Chardi, Hikka, Kasa, Karmiroga, Netraroga. cold, headache, cough, influenza, earache, fever, colic pain, sore throat, bronchitis, asthma, hepatic diseases, (15)(16)

#### **Dose**

1-3 ml of the drug in juice form.

1-2 g of the drug in powder form (seed). (15)

#### • Indian medicinal plant

In this book botanical source, synonyms, description, verities, and therapeutic uses are mention.

#### • Dravyaguna vigjan

In this book this botanical source, gana, synonyms, description, verities, constituents, action on different systems of body, properties and action, formulations, therapeutic uses, part use, dose is mention. (18)

#### • Sushruta samhita

In sushruta samhita mention in the sutra sansthan under the sursadi gana<sup>(20)</sup>

#### • Major herbs of Ayurveda

In this book botanical source, habitat, botanical description, traditional uses, ethnoveterinary usages, chemical constituents, pharmacological activities like immunomodulatory activity, antistress, antimicrobial, anti-inflammatory, antiasthmatic, anti-cancer, hypotensive activity, analgesic activities are mention. (21)

#### 4.3.2 Eranda Karkati<sup>(22)</sup>

**Botanical Name-** *Carica papaya* 

Family- Caricaceae

Vernacular name:

Language	Vernacular name
Hindi name	Papita
English name	Papaya fruit
Bengali Name	Papeya
Kannada Name	Parangimara
Telugu name	Boppayi
Tamil Name	Pappali

#### **Synonyms**

Nalaparni- The stalk of leaves are cylindrical.

MadhuKarkati, Erandachirbhata

#### Habitat

Papaya plant is a native of West Indies and Central America. It is now cultivated all over India in moderate climate. It is a small tree growing to a height of 4-6 m. The ripened papaya fruit is well known for its health benefits.

#### **Properties**

Characters	Properties
Rasa	Katu (Pungent), Tikta (Bitter)
Guna	Laghu (Light for digestion), Ruksa (Dry), Tikshna (Pungent)
Vipaka	Katu (Undergoes to pungent taste conversion after digestion)
Veerya	Ushna (Hot)
Karma	Kaphavatahara (pacifies Kapha and Vatadosha), Hridya( Benefits the heart)

Part used- Fruit, Leaf, Latex, Seed

**Dosage-** Leaf infusion 40-80 ml

Seed powder- 0.5- 1g

**Latex-** 3-6 g

#### **Chemical composition:**

The green parts of the plant and seeds of the papaya contain an alkaloid carpaine. The seeds also contain carpasamine. The latex of the plant contains papain, chymopapain and pseudocarpaine. The ripened fruit of papaya contains vitamin A, thiamine, riboflavin, niyacin and vitamin C. The latex of the unripe fruit is collected by giving a cut and the collected latex is dried under shade. papaya contains the digestive enzyme, papain. It is used in treating sports injuries, other causes of trauma, and allergies. Being rich in Vitamin A, pro-vitamin A carotenoid phytonutrients and C, Papaya fruit is an excellent anti-oxidant. (22)(23)

#### Uses

The milk from the raw papaya is applied over mouth ulcer to treat the milk of the raw papaya is applied over insect bites, scorpion bitten area to relieve the pain. The cold infusion prepared from the leaves of papaya is given in a dose of 20-30 ml to treat local swelling, fever, difficulty during micturition. The ripened fruit of papaya is consumed to relieve constipation. The raw papaya is consumed as vegetable to treat intestinal worm, piles and hepatomegaly. The milk from raw papaya is applied over wounds and bleeding piles to treat it. The leaves of the papaya plant are slightly heated and applied locally over the part of the body to relieve pain and swelling. The milk of raw papaya is applied locally to relieve skin diseases like eczema, ringworm. The papaya fruit is consumed to strengthen the muscles of the heart. (22)(23)

#### **Ayurvedic Indications**

Kasa, Shwasa, Yakrutodara, Pleehodara, Mandagni, Shotha, Visha.

#### • Drvyaguna vijyan

In this book this botanical source, gana, synonyms, description, verities, constituents, action on different systems of body, properties and action, formulations, therapeutic uses, part use, dose is mention. (24)

#### • Major herbs of Ayurveda:

In this book botanical source, habitat, botanical description, traditional uses, ethnoveterinary usages, chemical constituents, pharmacological activities like antifertility, diuretic activity, hypotensive, hypolipidaemic, anti ulcerogenic, anthelmintic, wound healing, antiabiotic, antimicrobial and safety profile and dose is mention. (25)

#### • Indian medicinal plants:

In this book botanical source, synonyms, distribution, botanical description, part uses, properties and uses are mention

#### 4.4 Review of formulation

#### 4.4.1 Phytosome

Preparation of medicine from the different parts of the herbal material is followed from the ancient time. Today is time of phyto medicine and phytosome is one of them which are a novel drug delivery system in herbal medicine field. The term "phyto" means plant while "some" means cell-like. (26) Phytosomes forms a bridge between the convectional delivery system and novel delivery system. Phytosomes show better stability as chemical bond is formed between phsospholipid molecule and phytoconstituent. It enhances the absorption of lipid insoluble polar phytoconstituents through oral as well as topical route showing better bioavailability, reduce dose. (27)(28)

# Advantages over the conventional dosage form $^{(29)}$

- 1. Phytosome are better bioavailable botanical extracts, dramatically enhance bioavailability due to their complex with phospholipids and delivers faster and improved absorption in intestinal tract.
- 2. Phytosome permeates the nonlipophillic botanical extract to be better absorbed in intestinal lumen.
- 3. Phytosome will be given in small quantity and desired results can be achieved.
- 4. Phytosome is widely used in cosmetics due to there more skin penetration and have a high lipid profile.
- 5. Phytosome are been used to give liver protectant flavonoids because they were easily bioavailable.

# EQUIPMENT, MATERIAL, AND EXPPERIMENTAL SETUP

# 5.1. List of Equipment

S. No.	Equipment
1.	Digital pH meter
2.	Hot Plate
3.	Hot Air over
4.	Muffler Furnish
5.	UV spectrophotometry
6.	Disintegration apparatus
7.	Dissolution apparatus
8.	Roche Friabilator apparatus
9.	Monsanto hardness tester
10.	Compound microscope
11.	TEM
12.	SEM
13.	Electron Micriscope
14.	Abbe's refractometer

# **5.2.** List of Chemicals

Ethanol	Dragndrofft reagent	Iodine	Benedict reagent
Methanol	Wagner reagent	Fehling B	Biurit's reagent
Hydrochloric acid Faric chloride		Fehling A	Dicloromethene

Mayer reagent	Lead acetate solution	Sudan 3	Soya lecithin
Salphuric acid	Acetone	Hexane	Sodium Hydrooxide
Pyrimidine	Acetic acid	N-hexane	Potassium
			Dihydrogen
			Orthophosphete
Toluene	Copper acetate	Sodium carbonate	Sodium nitripros
Ethyleacetate	Slica gel G	Sodium citrate	

# 5.3.List of drugs

- Tulsi
- Papaya leaves

#### RESEARCH METHODOLOGY

- Procurement of raw herbs from the local market of Jalandhar (Punjab)
- Authentication of raw herbs
- To study the classical and recent literature review regarding the disease and herbs used in formulation
- ➤ Pharmacognostic and phytochemical study of raw herbs
- Microscopic and macroscopic study of raw herbs.
- Phytochemical investigation of raw herbs.
- Physicochemical analysis of raw herbs
- 1. LOD at 110<sup>0</sup>C
- 2. FM
- 3. Total ash at  $450^{\circ}$ C
- 4. Acid insoluble ash
- 5. Water soluble extractive
- 6. Alcohol soluble extractive
- 7. Prepare the different dosage form
- Evaluation of prepared formulation
- Tablets
- a) Size and shape
- b) Appearance
- c) Uniformity of weight
- d) Disintegration test
- e) Dissolution test
- f) Mechanical strength
- g) Friabilty test
- h) Thin layer chromatography TLC
- i) High performance thin layer chromatography HPTLC
- Capsule
- a) Uniformity of weight

- b) Disintegration test
- c) Dissolution test
- Phytosome
- a) Determination of % yield
- b) Determination of particle size
- c) Determination of entrapment efficiency
- d) Determination of drug content
- e) Scanning electron microscopy (SEM)
- f) Fourier-transform infrared spectroscopy (FTIR)
- g) pH measurement
- h) Thin layer chromatography(TLC)
- i) High performance thin layer chromatography (HPTLC)
- Syrup
- a) Description, Colour
- b) Odour
- c) Total ash
- d) Acid insoluble ash
- e) Water-soluble extractive
- f) Alcohol soluble extractive
- g) PH
- h) Total sugar content
- i) Viscosity
- j) Thin layer chromatography (TLC)
- k) High performance thin layer chromatography( HPTLC)
- 1) Test for heavy metals
- m) Phytochemical analysis of formulation.
- > Stability study of prepared dosage form.
- > Comparative study of prepared formulation.

# Chapter 7 EXPECTED OUTCOMS

Dengue is the most rapidly spreading mosquito-borne viral disease in the world. There are not specific treatment of this is available .five type of dengue virus are founded these are DENV-1,DENV-2,DENV-3, DENV-4 and DENV-5 which is discover in the 2014. Only symptomatic treatment of it is available and different drug is used for different symptom.. In this study we are trying overcome these problems and One of the ingredient of C-pills papaya leaves have bitter taste by preparing different dosage forms of C-pills like phytosomes, syrup and capsule. Phytosomes are noval drug delivery system and that could be a promising new therapeutic approach for the treatment by the herbal drugs. Phytosome are better bioavailable botanical extracts, dramatically enhance bioavailability due to their complex with phospholipids and delivers faster and improved absorption in intestinal tract. During this study we try to make C-pills more effective by the addition of some other herbs in to it.

# PURPOSED WORK PLAN AND TIME LINES

	PURPOSED WORK PLAN AND TIME LINES																		
	Month List																		
Sr. no	Work List	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18
1	Indentification of topic																		
2	Review																		
3	Procurement of raw material																		
4	Authentication of raw material																		
	pharmacognostic &																		
5	phytochemical study																		
6	Perpration of formulation																		
7	Evaluation of prepared formulation																		
8	Stability study of prepared dosage form																		
	Data interpetation & Report writing																		

# **Result & discussion**

- 1. Authentication of raw herbs
- 2. Procurement of raw herbs
- 3. Macroscopic characters of (Tulsi)

S.no	Characters	Tulsi leaf	Papaya leaf
1	Colour	Green	Green
2	Odour	Aromatic	Characterstic
3	Taste	Characteristic	Better
4	Size	2.5-5 cm long, 1.6-3.2 cm wide	15 -25 cm diameter
5	Shape	elliptic-oblong	Palmate
6	Surface	Smooth at dosel surface	Smooth
7	Features	Hairs present on ventrel surface	5-10 cm long stalk is present

## 4. Phytochemical parameters

Sr.No	Paremeters	Drugs	Sample 1	Sample	Sample3	Average	Standard
				2			
1	Foreign	Tulsi	1.4%	2.1%	1.9%	1.8%	NMT(32)
	matter						2%
		Papaya	3.2%	2.8%	5.2%	3.7%	-
2	Loss on	Tulsi	7.1%	8.4%	7.9%	7.8%	
	drying	Papaya	11.5%	10.8%	11.9%	11.4%	9.57% <sup>(30)</sup>
3	Total ash	Tulsi	13%	10%	12.5%	11.83%	NMT <sup>(32)</sup>
							19%
		Papaya	13.5%	14%	13%	13.5%	12.7% (30)

4	Acid	Tulsi	2%	2%	1.5%	1.83%	NMT
	insoluble						3% <sup>(32)</sup>
	ash	Papaya	2%	2.5%	2.5%	2.3%	1.75% (30)
5	Alcohol	Tulsi	7.2%	4.8%	7.2%	6.4%	NLT
	soluble						6% <sup>(32)</sup>
	extractive	Papaya	10.4%	13.6%	9.6%	11.2%	8.4% <sup>(31)</sup>
6	Water	Tulsi	11.2%	12.8%	15.2%	13.06%	NLT <sup>(30)</sup>
	soluble						13%
	extractive	Papaya	26.4%	31.2%	30.4%	29.3%	17.7% (31)

# 5. Phytochemical screening

S.	Name of	Name of test	Tulsi	Papaya
no	Phytoconstituents			
1	Alkaloids	(a)Mayer, s test	+	+
		(b) Wagner, s test	+	+
		(d) Dragendroff's	+	+
		Test		
2	Carbohydrates	(a) Molisch's Test	+	+
		(b) Benedict's Test	-	-
		(c) Fehling's Test	+	+
3	Glycosides	(a) Modified	-	-
		Borntrager's Test		
4	Flavonoid	(a) Alkaline reagent	+	+
		test		
		(c) ) NH4OH test	+	+
5	Tanin	(a)Farric chloride	+	+
	ı		I.	1

6	Phenolic compound	(a)Farric chloride	-	-
7	Steroid	(a)salkowski's	+	-

# Chapter 10 SUMMARY & CONCLUSION

Yet to be done

#### REFERENCES

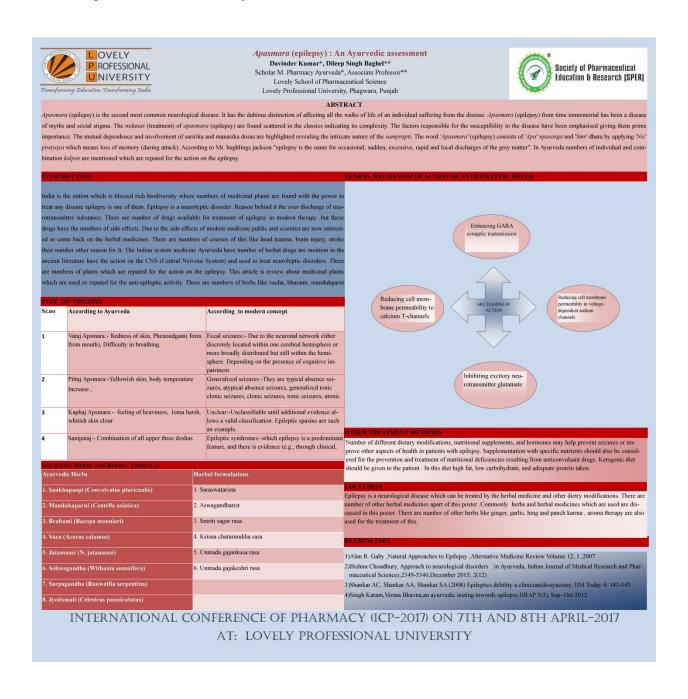
- 1.Andrew W. Taylor-Robinson, A Putative Fifth Serotype of Dengue Potential Implications for Diagnosis, Therapy and Vaccine Design, International Journal of Clinical & Medical Microbiology, 2016, 1: 101.
- 2. https://www.nhp.gov.in/ayurvedic-perspective-of-dengue-fever\_mtl
- 3. Janagal Bhageshwary ,et al, dengue in ayurvedic perspective and its management a Review article, International Ayurvedic Medical Journal ISSN: 2320 5091,
- 4. Acharya Shukal V, Ayurvediya Vikriti Vijyan, 17<sup>th</sup> Chepter, Chaukhmba Sanskrit Pratishthan, Delhi; 2004, p. 248
- 5. Dr. Kumar V, Roga Vijyan, Nath Pustak Bandhar, Rohatak,2<sup>nd</sup> part, 1<sup>st</sup>chapter, p. 20-21
- 6. Dr. Sumit Srivastava, an ayurvedic insight towards dengue fever, european journal of pharmaceutical and medical research, 2015;2(4): 693-698
- 7. Siti Latifah Abd Kadir, Harisun Yaakob, Razauden Mohamed Zulkifli, Potential anti-dengue medicinal plants: a review, J Nat Med, 12 March 2013.
- 8. Holger H et al, Serotype-specific Differences in Dengue Virus Non-structural Protein 5 Nuclear Localization, the journal of biological chemistry, 2013; VOL. 288, pp. 22621–22635
- 9. Sobia I and Usman A Ashfaq, A brief review on dengue molecular virology, diagnosis, treatment and prevalence in Pakistan, BioMed Central Ltd, 2012
- 10. JK Lalla et al, A Review on Dengue and Treatments, journal of pharmacology and toxicological studies,2014;vol 2; 4
- 11. Dengue guidelines for diagnosis, treatment, prevention and control, A joint publication of the World Health Organization and the Special Programme for Research and Training in Tropical Diseases,2009
- 12. Gupta N et al, Dengue in India, Indian J Med Res, 2012;136; 373-390
- 13. http://nvbdcp.gov.in/den-cd.html Date 18Aug2017
- 14. Battula P et al, Caripill: An Advance Treatment for Dengue, International journal of pharmacy & pharmaceutical research, 2016 Vol.:6, Issue:2

- 15. Anonymous, The Ayurvedic Pharmacopoeia of India, ministry of ayush ,Part 1, New Delhi.Vol.2,p 174
- 16. Verma S, Chemical constituents and pharmacological action of *Ocimum sanctum* (Indian holy basil-Tulsi), The Journal of Phytopharmacology 2016; 5(5): 205-207
- Lt-Colonel Kirtikar K.R, Indian Medicinal Plants, International Book Distributor, Dehradun;
   p. 1965
- 18. Prof. Sharma P.V, DravyaGuna Vijyan, Chaukhmba Bharti Academy, Varanasi, Vol 2; p. 516
- 19. Singh K et al, Prevention and control of dengue by herbal remedies, Journal of Chemical and Pharmaceutical Research, 2016; 8(3):708-71
- 20. Dr. Sharma A, Sushruta samhita, Chaukhmba Bharti Parkashan, Part 1, P 216
- 21. Major Herbs of Ayurveda, Compiled by the Dabur Reserch Foundation, Edited by Elizabeth M Williamsom, p. ,201
- 22. Yogiraj V et al, Carica papaya Linn: An Overview, International journal of herbal medicine, 2014; 2 (5): 01-08
- 23. Krishna KL et al, review on nutritional, medicinal and pharmacological properties of papaya, natural product radiance, 2008; vol 7: p 364-373
- 24. 17. Prof. Sharma P.V, DravyaGuna Vijyan, Chaukhmba Bharti Academy, vol 2, Varanasi; p.372
- 25. Major Herbs of Ayurveda, Compiled by the Dabur Reserch Foundation, Edited by Elizabeth M Williamsom, p.88
- 26. Shelke SS et al, Phytosomes A New Herbal Drug Delivery System, International Journal of Research in Pharmaceutical and Biomedical Sciences, 2012;vol 3:p 1709-1715
- 27. Amith Kumar B et al, Phytosomes as Novel Drug Delivery System for Herbal Medicine –A Review, Systematic Reviews in Pharmacy,2017;vol 8:p 5-7
- 28. Aniket et al, Preparation and Characterization of Soy-Phytosomes using Ethanol: Water (70:30) Solvent System, International Journal of Pharmaceutical Sciences, 2015; vol 1(1), 18-26
- 29. Gandhi A et al, Recent Trends of Phytosomes for Delivering Herbal Extract with Improved Bioavailability, Journal of Pharmacognosy and Phytochemistry, 2012; vol 1: p 6-14

- 30. Lalla J.K. and Ogale S, pharmacognistic evaluation of leaves of carica papaya linn, world journal of pharmacy and pharmaceutical sciences, 2015; vol 4:p1066-1081
- 31. Zunjar V et al, Pharmacognostic, Physicochemical and Phytochemical Studies on Carica papaya Linn. Leaves, Pharmacognosy Journal, 2011; vol 3:p5-8
- 32. Anonymous, The Ayurvedic Pharmacopoeia of India, ministry of ayush ,Part 1, New Delhi.Vol.2,p 175

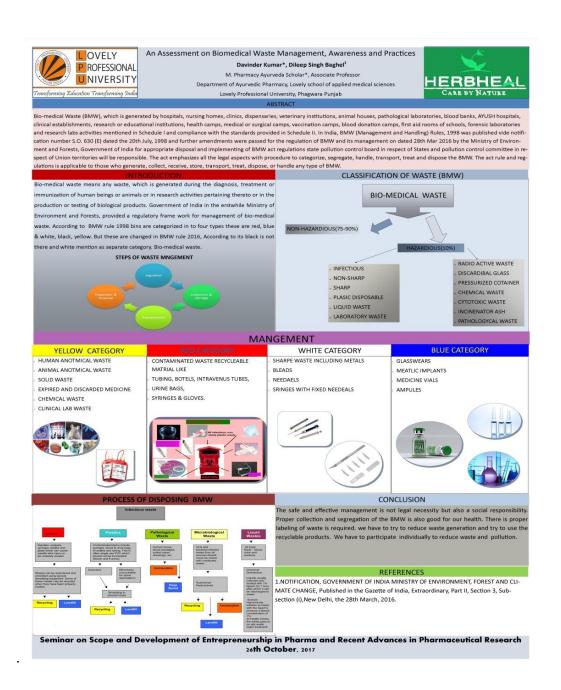
#### Appendices I

#### Present a poster in SPER( Society of Pharmaceutical Education & Resarch) ICP-2017 LPU



#### **Appendices II**

Present a poster in Seminar on Scope and Development of Entrepreneurship in Pharma and Recent Advances in Pharmaceutical Research 26th October, 2017, HHRC Amritsar



# Report

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#### TOPIC APPROVAL PERFORMA

LIT (Pharmacy)/Department of Pharmaceutical Sciences

Program: 5303H::B. Pharmacy (Ayurveda) – M. Pharmacy (Ayurveda)(Dual Degree)

COURSE CODE : APH625		REGULAR/BACKLOG:		OG: Regular	GROUP NUMBE		ER: PHRRGD0056		
Supervisor Na	sme: Dileep	Singh Baghel	UID: 1	5210		Desi	gnation:	Associate Professor	
Qualification :				Research Ex	perience :				
R.NO. NAME OF STUDENT  Davinder Kumar		RE	EGISTRATION NO	BATCH	SECTION	CONTACT NUMBER			
1	Davinder Kumar		11	1203537	2012	Y1653	9988954422		
SPECIALIZATION AREA : Ayurvedic Pharmacy Supervisor Signature:									
PROPOSED TO	OPIC:	AREA : Ayurvedic Pharmacy Supervisor Signature:							

	Qualitative Assessment of Proposed Topic by PAC					
Sr.No.	Parameter	Rating (out of 10)				
1	Project Novelty: Potential of the project to create new knowledge	6.50				
2	Project Feasibility: Project can be timely carried out in-house with low-cost and available resources in the University by the students.	7.00				
3	Project Academic Inputs: Project topic is relevant and makes extensive use of academic inputs in UG program and serves as a culminating effort for core study area of the degree program.	6.50				
4	Project Supervision: Project supervisor's is technically competent to guide students, resolve any issues, and impart necessary skills.	7.50				
5	Social Applicability: Project work intends to solve a practical problem.	7.00				
6	Future Scope: Project has potential to become basis of future research work, publication or patent.	6.75				

PAC Committee Members					
PAC Member 1 Name: Dr. Amit Mittal	UID: 13145	Recommended (Y/N): Yes			
PAC Member 2 Name: Saurabh Singh	UID: 12208	Recommended (Y/N): Yes			
PAC Member 3 Name: Dr. S. Tamilvanan	UID: 16391	Recommended (Y/N): Yes			
PAC Member 4 Name: Dr. Navneet Khurana	UID: 18252	Recommended (Y/N): Yes			
DAA Nominee Name: Dr. Sazal Patyar	UID: 17050	Recommended (Y/N): NA			

Final Topic Approved by PAC: Pharmaceutical standardization, product development and quality control aspect of C- Pills a marketed product

Overall Remarks: Approved

PAC CHAIRPERSON Name: 11045::Dr. Monica Gulati Approval Date: 25 Apr 2017