### CONTENT

- Introduction
- Causes
- Clinical Sign
- Types
- Screening Models



### INFLAMMATION

- Inflammation can be simply defined as a protective response of body's immune system against injury.
- It is a process by which our body's white blood cells protect us from infection with foreign particles.
- Inflammation is body's natural reaction against injury & infection.

### CAUSES OF INFLAMMATION

- There are various causes of inflammation as follows :
- Physical Agents : Heat, cold, Radiation
- Chemical Agents : Acid, base, toxic gases
- Biological Agents : Viral bacteria , fungal infections .
- Metabolic Products : Uric Acid , Urea .
- Circulation Disorder : Thrombosis , infarction

### CLINICAL SIGN OF INFLAMMATION

- Redness (rubor)
- Swelling (tumour)
- Heat (calor)
- Pain (dolor)
- Loss of function (Functio Laseo)



# TYPES OF INFLAMMATION

Depending on the time of duration , inflammation are of two types :

- Acute Inflammation
- Chronic Inflammation
- ACUTE INFLAMMATION : It occurs for a short time period and a rapid onset
- The characteristics of acute inflammation are:
- Accumulation of fluid & plasma protein on affected side
- Activation of Platelets & Neutrophills .

- **CHRONIC INFLAMMATION :** It occurs for a long time period & remain for longer duration
- The characteristics of chronic inflammations are :
- Mononuclear Infiltration
- Activation of Lymphocytes & Macrophages .

## SCREENING MODELS

#### **IN-VITRO METHODS**

- Cox Assay
- Mast cell degranulation
- Platelet-neutrophils adhesion
- Adhesion assays
- Inhibition of NO production induced by TNF-a in mouse macrophages

# IN-VIVO METHODS

- UV-B induced erythema in guniea pigs
- Carrageen induced paw edema model
- Croton oil induced ear edema in mice
- Plural exudation method
- Adjuvant arthritis
- Air pouch model
- Cotton pellet induced granuloma

#### CARRAGEENAN INDUCED PAW EDEMA IN RAT

- **PRINCIPLE :** It is based on the principle that the ability of test drug to inhibit edema produced on the hind limbs of the experimental animals after the injection of inflammatory agent
- ANIMAL : Male wistar / Albino rat {100-150gm}
- **INDUCING CHEMICAL :** carageenan
- **VOL. OF ADMINISTER** : 0.01ml by orally or s.c. route
- STD. Drug Use: Diclofenac

## PROCEDURE

- > Animals are divided into three groups . Starved overnight with appropriate water and diet .
- > All groups are penetrated by inflammatory agents Eg. Careageenan before 30min. of testing in left paw hind limb of each animal
- The Control group receives distilled water only, while other receives test and standard drug respectively.
- The paw is marked with ink and immersed in water cell of plethismometer
- After 15,30 and 45 min. of injection paw edema volume of test and standard is compared with control group.





: Before carrageenan injection



B: After carrageenan injection

#### CROTON OIL EAR EDEMA

- **PRINCIPLE :** It is based on the principle that the croton oil induces inflammation and edema at applied site in animals and local anti-inflammatory agent can be evaluated in this model .
- ANIMALS : Mice (22-30gm) Rat (75-80gm) INDUCING CHEMICALS :
- For mice Croton oil : ethanol : pyridine : ether (1:10:20:69)
- **For rat** Croton oil : ethanol : pyridine : ether (1:10:20:66)
- VOL. OF INJECTION : 0.01ML

#### PROCEDURE

- The Std test compound dissolve in above solution and applied on ear of all group of animal
- Fest compound were dissolved in irritant solution in conc.0.03 to 1 mg/ml for mice and 3 to 10 times higher conc. For rat
- > Applied on both side of right ear 0.01ml in mice and 0.02ml in rats of all group while control group receives only vehicle solution
- After 4hrs of the application animals are scarified and ear is removed and disc of 8mm is obtained by cutting the ear.

- **OBSERVATION :** The disc are weight and difference between test std. and control is compared .
- The difference in the weight is indication of the antiinflammatory activity .
- % Anti-inflammatory = wt. of wt. of Effect treated contralater \* 100 ear control ear

wt. of contralateral control

### ANTI-PYRETIC DRUGS

- The drugs which lowers the elevated body temperature to the normal.
- Drugs used are :
- > Aspirin
- > Paracetamol (PCM)



# SIGN AND SYMPTOMS

- Nausea
- Vomiting
- Jaundice
- Abdominal pain
- Feeling and being sick
- Stomach pain
- Headache
- High temperature
- Loss of appetite
- Chills (Shivering)

### SCREENING MODELS

Brewer's yeast suspension method
Antipyretic activity in Rabbits
Antipyretic activity in Rats



### BREWER'S YEAST SUSPENSION METHOD

- Animal : Male or Female wistar rat (150gm)
- Inducing agent : Brewer's Yeast
- Std. Drug : Phenacetin 100mg/kg orally
- Procedure
- > 15% suspension of brewer's yeast in 0.09% saline is prepared
- Group of 6 male and female wistar rat with body weight of 150gm are used .
- Insert thermocouple to the depth 2cm into the rectum of the rat and record initial temp.

- > 10mg/kg brewer's yeast suspension inserted by s.c route in the back below the nape of the neck.
- Massaged at the site of injection in order to spread the suspension under the skin and withdrawn the food after yeast administration.
- > After, 18 hours the administration of the yeast the rise in rectal temp. recorded the measurement repeat after 30 min .

- > Only animal with body temp. of at least 38\*c are taken to the test .
- > Animal receive test compound or standard compound by oral administration
- Rectal temperature are recorded again 30,60,120 and 180 min post dosing
- Evaluation
- Difference between initial temp. and after dosing temp.
- Maximum reduction in rectal temp. is compare to the control group is calculated .

#### ANTI-PYRETIC TESTING IN RABBIT

- Animal : Male or female Rabbit (3-5kg)
- Inducing agent : Lipopolysaccharides
- Std. Drug : Indomethacin 2.5mg/kg
- Procedure
- > Lipopolysacchrides from gram negative bacteria eg. E-coli induced fever in rabbit .
- Rabbit of both sex of the various strains with body weight between 3-5 kg are used .
- > Animal placed in suitable cage and introduced a thermocouples into the rectum

Of the rabbit and allow to adapt to the cage for 60min

- > 0.2 microgrm lipopolysacchride are injected i.v. at the dose 0.2ml/kg body weight into the ear.
- > 60 min later test compound administered either sc or by oral route and body temp. are monitored for at least 3 hrs
- > decreased of the body temp. atleast 0.5\* c for more than 30min found as positive effect.