Human Journals

Research Article
February 2021 Vol.:20, Issue:3

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Evaluation of Anti-Pyretic Activity of *Woodfordia fruticosa* Using Brewer's Yeast Induced Pyrexia in Albino Rats



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Submitted: 03 January 2021
Revised: 23 January 2021
Accepted: 12 February 2021



www.ijppr.humanjournals.com

Keywords: Anti-pyretic activity, *Woodfordia fruticosa* (Dhataki), Paracetamol, Yeast, albino rats.

ABSTRACT

Background: Many herbal plants are having antipyretic effects. W. fruticosa traditionally used to treat fever pain and inflammation. Aim: The present study investigates the antipyretic activity of the brownish extracts of Woodfordia fruticosa flower on brewer's yeast induced fever in experimental rats. Methods: 30 albino rats weighing 150-180g were used for study. Brewer's-yeast-induced pyrexia was used to investigate the antipyretic effects. 10ml/Kg (subcutaneous) of 0.5(w/v)suspension of brewer's Carboxymethylcellulose was injected to induce fever in all the experimental animals. After 18hrs, the rectal temperature was taken and the animals were administered Woodfordia fruticosa (100mg/kg, 200mg/kg, and 400mg/kg) and paracetamol (standard group, 30mg/kg) orally. Result: The body temperature of the rats was measured rectally over a period of 3hours. W. fruticosa (200mg/kg and 400mg/kg) significantly reduced yeast-induced pyrexia when compared with the standard drug. Thus, this experiment shows that the antipyretic effect of W. fruticosais dose-dependent and the effect is as a result of the flavonoid component of the extract. Conclusion: These data therefore suggest that extract of W. fruticosa possesses significant antipyretic activity and it mechanism could be by inhibition of release inflammatory mediators and prostaglandins.

INTRODUCTION:

Over the centuries, plants have been used for their medicinal values. However, many plants have yet to be studied and relatively few scientific researches have been carried out.^{1, 2} Therefore, assessment of medicinal plants is of special interest because of rich heritage and the continuous use among large portion of people.¹⁻³

Fever, is known as pyrexia and may occur due to infection, inflammation, or any tissue damage and disease states. It is defined as the elevation of core body temperature above normal; in normal adults, the average oral temperature is 37°C (98.6°F) (1). Fever is a medical sign rather than a disease and it may be caused as a secondary impact of infection, malignancy or other diseased states. Fever occurs when the body's thermostat (located in the hypothalamus-anterior pituitary) resets at a higher temperature, primarily in response to an infection. From the recent scientific discovery, most of the antipyretic drugs have been developed to reduce elevated body temperature, of which many acts by the mechanism of inhibition of the COX-2 expression to reduce PGE2 biosynthesis.⁴⁻⁷

It is currently accepted that prostaglandin E2 (PGE2) is the final fever mediator in the brain, specifically in the pre-optic area of the anterior hypothalamus. Many drugs on chronic usage can able to result in several side effects including gastrointestinal, renal, hepatic, central nervous system and dermatological effects. A rich heritage of knowledge on preventive and curative medicines was available in ancient scholastic work.^{4, 5, 7}The beneficial medicinal effects of plant materials typically result from the combinations of secondary products present in the plant. Many of the plants were found to show the antipyretic activity hence the use of traditional plants as relieving agents can be advantageous in many aspects such as easy availability, economical with no or limited side effects.⁸

MATERIALS AND METHODS:

Woodfordia fruticosa, also called Dhai, Dhataki, Fire Flame Bush is a species of plant from the Lythraceae family. In Ayurveda this herb is used to treat many diseases. Its flower is analgesic antipyretic and astringent due to pitta dosha. It is widely cultivated in tropical regions, usually between 0 and 300 m altitude, and grows well in brushwood, bush margins, grassy fields, and secondary forest, seasonally flooded grassy fields and sandy coasts and along roadsides and riverbanks, although the species cannot tolerate excessive water or heat. 9-17

Citation: Kapil Jangir et al. Ijppr.Human, 2021; Vol. 20 (3): 259-266.

Plant Taxonomy: 18-21

Table No. 1: Scientific classification of Woodfordia fruticosa

Cynonyma	Dhai, Dhati, Dhataki,				
Synonyms	Dawai, Dhavi				
Kingdom	Plantae				
Subkingdom:	Tracheobionta				
Division:	Magnoliophyta				
Class:	Magnoliopsida				
Subclass:	Rosidae				
Order:	Myrtales				
Family:	Lythraceae				
Genus:	Woodfordia				
Species:	fruticosa				
icals:	()				

Drugs and chemicals:

Paracetamol was purchased from CDFCL. The solvents and other chemicals of analytical grade were used and obtained from the institute's central store. Brewer's yeast was purchased from local market.

Collection& extraction of Flowers:

The fresh flowers of W. *fruticosa* were collected from the local areas of Alwar district and are air dried under shade for two days and then coarsely powdered.

Defattation of powdered Woodfordia fruticosa flower

200 g coarse flower powdered were defatted with 400 ml petroleum ether (40-60°C) using soxhlet apparatus. Extraction was continued until a drop of solvent from siphon tube, when evaporated on filter paper, did not leave a greasy spot (approximately 10-12 cycles). After the defattation, mark was taken out from extractor and spreaded as a bed on a clean paper and dried till evaporation of petroleum ether.²²⁻²⁴

Ethanolic extraction of W. fruticosa:

The dried mark obtained after defattation was packed in soxhlet apparatus and extracted with 400 ml ethyl alcohol in soxhlet apparatus. Extraction was continued until a drop of solvent from the siphon tube, when taken on TLC plate and sprayed with concentrated sulphuric acid, does not give a black spot. Brownish extract thus obtained, was collected and solvent was evaporated under reduced pressure. The percentage yield of ethanolic extract was found to be 10.12 % which is comparable with the reported standard extractive value.²⁵⁻²⁷

Animals

30 albino rats weighing between 150-180g of either sex were used for this experimental study. They were divided into five groups (n=6). To the rats laboratory, diet and water *ad libitum was provided and* were kept in polypropylene cages. Which are kept in a constant humidity (55%), temperature at (22± 2°C), and exposed to dark and light {12hr} every day the bedding materials of the cages were changed. The experimental protocol was approved by IAEC, with proposal no APC/CPCSEA/2020/1709, for using animals in this experiment. Animals were fasted overnight with free access to water before each experiment.^{24, 28}

Table No. 2: Grouping and dose of drug for anti-pyretic activity

Group	Drug and Doses					
Brewer's yeast-induced pyrexia						
Group I Normal Control (Received 0.5% CM						
Group II	W. fruticosa100mg (i.p.)					
Group III	W. fruticosa200mg (i.p.)					
Group IV	W. fruticosa400mg (i.p.)					
Group V	Paracetamol 30 mg/kg (oral)					

Induction of Pyrexia:

Fever was induced by injecting 10ml/Kg (subcutaneous) of 0.5(w/v) suspension of brewer's yeast in Carboxy methylcellulose below the nape of the neck. The temperature was measured after 18hours using rectal thermometer.^{29, 30}

Drug administration:

After 18 h of yeast injection 0.5% CMC was administered orally to the control Group I. The extracts of *W. fruticosa* were administered at doses of (100, 200 and 400 mg/kg b.w. p.o) to groups II, III, IV animals respectively. Group V animals received the standard drug Paracetamol (30 mg/kg b.w. p.o) and Rats were restrained for recording rectal temperature by at the 18 h, immediately before extracts, normal saline or paracetamol administration, and again at one-hour intervals up to the 21 h after yeast injection.

Statistical Analysis:

Results are expressed as mean \pm SEM. Statistical analysis of data was performed using ANOVA to study differences among the means.

RESULTS AND DISCUSSION:

Fever is known to be caused by several endogenous pyrogens such as interleukin-1 β , interleukin-6, interleukin-8, tumor necrosis factor- α , macrophageprotein-1 and prostaglandins. Prostaglandin synthesis may be activated by tumor necrosis factor- α and phospholipase A2. Brewer's yeast induces both TNF- α and prostaglandin synthesis.

Table No. 3: Antipyretic Effect of Ethanolic extract of *Woodfordia fruticosa* on yeast induced pyrexia in rats.

	Dose mg/kg	Rectal temp. °C before and after treatment							
Treatment		Normal		30 min after treatment	60 min after treatment	90 min after treatment	120 min after treatment	180 min after treatment	
Control	-	37.7±0.18	38.7±0.34	38.6±0.21	38.5±0.04	38.5±0.24	38.5±0.12	38.5±0.16	
Paracetam ol	30	37.6±0.12	38.6±0.24	38.2±0.13	37.8±0.09	37.7±0.04	37.6±0.26	37.62±0.06	
EEWF	100	37.7±0.17	38.6±0.32	38.2±0.04	38.12±0.05	37.98±0.16	37.90±0.07	37.82±0.05	
EEWF	200	37.6±0.21	38.6±0.18	38.12±0.05	37.9±0.05	37.82±0.04	37.7±0.04	37.7±0.06	
EEWF	400	37.6±0.08	38.6±0.15	37.9±0.06**	37.8±0.03**	37.8±0.06**	37.68±0.08**	37.67±0.07	

Values are expressed as mean \pm SEM (n=6). **P<0.01 Statistically.

The subcutaneous injection of yeast suspension (Nape of neck) markedly elevated the rectal temperature after 18 h of administration. Treatment with the WF at doses 200 and 400 mg/kg significantly (p<0.01) decreased the rectal temperature of the rats in dose dependent manner up to 3 h after pyrexia when compared with control (Table 3). Effect of 100mg/kg extract was negligible. The antipyretic effect was sustained throughout the remaining period of the experiment in a manner similar to the reference drug Paracetamol 30mg/kg, b.w. p.o.

The extracts are likely to reduce fever by reducing brain concentration of prostaglandin E2 especially in the hypothalamus through its action on COX-2 or by enhancement of the production of the body's own antipyretic substances.

The efficacy of the antipyretic effect of dhataki extract was observed to have increased with increased concentration (dose-dependent manner). This can be said to be due to the increased concentration of the component of the extract exhibiting antipyretic effects.

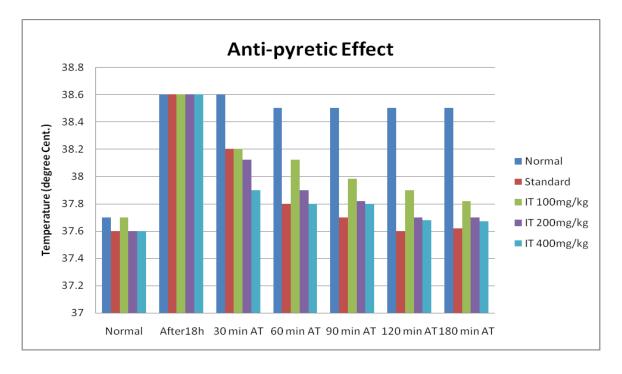


Figure No. 1: Graphical Representation of Anti-pyretic Effect of Ethanolic extract of *W. fruticosa* on yeast induced pyrexia in rats

CONCLUSION:

Antipyretic activity of bioflavonoid from *Woodfordia fruticosa* plant extract was confirmed by the present experimental studies. *W. fruticosa* extract with two different doses of 200mg/kg and 400mg/kg showed the reduced temperature levels in the wistar albino

experimental rats when compared to the temperatures of rats of control group. It is through the inhibition of prostaglandin (PGE2). From these results, it can be concluded that the *W. fruticosa* plant extract at the dose level of 200mg/kg has shown a marked decrease in temperature to normal. Hence possess antipyretic activity which was comparable with the paracetamol effect. Plant test extract of 100mg/kg was not so efficient as the dose level of 200mg/kg.

CONFLICT OF INTERESTS:

The authors declare that there is no conflict of interests regarding the publication of this paper.

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