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## FORMULATION DEVELOPMENT AND SUSTAINED RELEASE OF MATRIX TABLETS OF ANTI-ANGINAL DRUGS

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## ABSTRACT

The present investigation revealed that *Tamarindus kernels* mucilage and guar gum, Ethyl cellulose and HPMC appears to be suitable for use as a release retarding agents in the formulation of sustained release matrix tablets is good swelling, good flow and suitability for matrix formulation. From the dissolution study, it was concluded that *Tamarindus kernals* mucilage can be used as an excipient for making sustained release matrix tablets of Nicorandil. The formulated tablet was characterized by physical and chemical parameters, the *in-vitro* release of rate profile should the higher concentration of F2 polymer in tablet, the combination of hydrophilic and hydrophobic combination showed less result than use of alone.

## KEYWORDS

Angina, Nicorandil matrix tablet, HPMC and Ethyl cellouse etc.

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## **INTRODUCTION**

Tablet is a solid dosage form containing a unit dose of one or more medicaments. Tablets are flat or biconvex discs prepared by compressing a drug or a mixture of drugs with or without suitable excipients. Tablet may vary in shape and differ greatly in size and weight depending on the amount of medicinal substance and the intended mode of administration, with advancement in technology. That is convenient for patients perspective and utilize an approach that is unlikely to add complexity during regulatory approval process. To understand each dosage form, tablets here are classified by their route of administration and by the September – October 291 type of drug delivery system they represent within the route.

# EXPERIMENTAL WORK

#### Extraction of mucilage

*Tamarindus* seeds of 1kg are collected and washed with water to remove extra pulp.

These *tamarindus kernels* are then heated in a container along with sand for sometime, which gives brittleness to the external coat.

These heated kernels are crushed and removed the external coat to get the seeds.

The seeds obtained are kept in water for overnight and then boil for 5 to 7 hours, and then filtered to remove seeds.

Then the mucilage solution (filtrate) is heated for sometimes and kept in hot air oven at the temperature between 50 to 60°C until complete moisture is evaporated.

The dried mucilage is powdered and passed through sieve.no.100. Then the mucilage of 14gms is collected from 1kg of seeds and then stored well in an air tight container.

### **Evaluation tests for mucilages**

Treat the test solution with ruthenium red solution pink colour is obtained.

Treat the test solution with thionine solution after 15 minutes wash with alcohol.

Mucilage forms violet red colour.

Treat the test solution with Chinese ink, transparent spherical dilated fragments on blackbackground are observed.

#### **Formulation batches**

Take 10ml of aqueous solution and add 25ml of absolute alcohol with constant stirring. Aprecipitate is formed which is dried in air and examined for its swelling properties.

## **Tablet preparation**

All the ingredients were weighed accurately using digital weighing balance and werepassed through a 100 sieve.

The binder solution was prepared by dissolving *tamarindus kernels* mucilage in hexaneas solvent.

All the excipients are blended together. Then all the excipients, including drug were mixed in the binder solution according to their respective formulations (F1, F2, F3, F4 and F5).

The powder mixture was kept in hot air oven at 60°C for complete evaporation ofhexane.

The tablets were prepared by direct compression method by using 15mm biconvex punches at the pressure of 35psi.

#### DISCUSSION

The formulation development of Nicorandil sustained release matrix tablets was done with different concentrations of natural polymers as a binding agent. Various evaluation tests were conducted for the formulated batches. *Tamarindus kernels* mucilage has been checked for some evaluation parameters to know the flow properties, Bulk density, Tapped density, Compressibility index and Hausner's ratio are calculated to know the flow properties of the mucilage.

The released profile of all the formulation is shown in the Figure No.7. The sustained release matrix tablets of Nicorandil prepared with different natural polymers in various concentrations.

The order of cumulative percentage drug released was F4>F2>F5>F3>F1 with values of 74%, 73%, 64%, 57%, 52 % respectively. The present study revealed that the formulation F4 with Drug: Gum ratio of (1:3.6) should greater cumulate percentage drug release and the formulation F1 with Drug: Gum ratio of (1:1.8) should lower cumulative percentage drug release.

|      | Table No.1        |      |      |           |      |      |  |  |  |
|------|-------------------|------|------|-----------|------|------|--|--|--|
| S.No | Ingredients       | F1   | F2   | <b>F3</b> | F4   | F5   |  |  |  |
| 1    | Nicorandil        | 80   | 80   | 80        | 80   | 80   |  |  |  |
| 2    | Guar gum          | 320  |      |           | 150  |      |  |  |  |
| 3    | Tamarindus gum    |      | 320  |           | 150  | 150  |  |  |  |
| 4    | HPMC              |      |      | 320       |      | 150  |  |  |  |
| 5    | Ethyl cellulose   |      |      |           | 20   | 20   |  |  |  |
| 6    | Magnesium sterate | 2.5  | 2.5  | 2.5       | 2.5  | 2.5  |  |  |  |
| 7    | Talc              | 97.5 | 97.5 | 97.5      | 97.5 | 97.5 |  |  |  |
| 8    | Total weight      | 500  | 500  | 500       | 500  | 500  |  |  |  |

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# Physical properties of matrix tablets

#### Table No.2

| Formulation | Weight variation (%) | Thickness(mm) | Hardness(kg/cm2) | Friability(%) |  |  |  |  |
|-------------|----------------------|---------------|------------------|---------------|--|--|--|--|
| F-1         | 1.9                  | 4.6           | 4.1              | 0.50          |  |  |  |  |
| F-2         | 3.7                  | 4.7           | 3.9              | 0.45          |  |  |  |  |
| F-3         | 4.6                  | 4.7           | 4.8              | 0.35          |  |  |  |  |
| F-4         | 2.4                  | 4.7           | 4.9              | 0.78          |  |  |  |  |
| F-5         | 3.5                  | 4.6           | 4.8              | 0.65          |  |  |  |  |

## STANDARD ABSORBANCE OF NICORANDIL

#### Table No.3

| S.No | Concentration (µg/ml) | Absorbance at 260nm |  |  |
|------|-----------------------|---------------------|--|--|
| 1    | 0                     | 0                   |  |  |
| 2    | 10                    | 0.105               |  |  |
| 3    | 20                    | 0.165               |  |  |
| 4    | 40                    | 0.324               |  |  |
| 5    | 60                    | 0.459               |  |  |
| 6    | 80                    | 0.632               |  |  |
| 7    | 100                   | 0.765               |  |  |

### Table No.4

| S.No | Time<br>(Hrs) | Absorbance<br>(nm) | Concentration<br>(µg/ml) (X) | X*900/1000 | 5*X/1000 | Cumulative<br>Amount | %Cumulative<br>drug release |  |  |
|------|---------------|--------------------|------------------------------|------------|----------|----------------------|-----------------------------|--|--|
| 1    | 0             | 0                  | 0                            | 0          | 0        | 0                    | 0                           |  |  |
| 2    | 0.5           | 0.123              | 14.320                       | 0.159      | 7.160    | 5.433                | 5.433                       |  |  |
| 3    | 1             | 0.189              | 23.044                       | 20.739     | 0.115    | 60.274               | 60.274                      |  |  |
| 4    | 2             | 0.236              | 29.253                       | 26.325     | 0.146    | 20.885               | 20.885                      |  |  |
| 5    | 4             | 0.453              | 57.919                       | 52.127     | 0.289    | 26.614               | 26.614                      |  |  |
| 6    | 8             | 0.337              | 42.595                       | 38.335     | 0.212    | 52.339               | 52.339                      |  |  |
| 7    | 16            | 0.241              | 29.914                       | 26.922     | 0.149    | 38.484               | 38.484                      |  |  |
| 8    | 24            | 0.102              | 11.552                       | 10.395     | 0.057    | 26.979               | 26.979                      |  |  |

| S.No       | Time<br>(Hrs) | Absorbance<br>(nm) | Concentration<br>(µg/ml) (X) | X*900/1000 | 5*X/1000 | Cumulative<br>Amount | %Cumulative<br>drug release |  |
|------------|---------------|--------------------|------------------------------|------------|----------|----------------------|-----------------------------|--|
| 1          | 0             | 0                  | 0                            | 0          | 0        | 0                    | 0                           |  |
| 2          | 0.5           | 0.229              | 28.328                       | 25.495     | 0.414    | 10.414               | 10.414                      |  |
| 3          | 1             | 0.285              | 35.601                       | 32.040     | 0.178    | 25.673               | 25.673                      |  |
| 4          | 2             | 0.304              | 38.236                       | 34.412     | 1.191    | 32.321               | 32.321                      |  |
| 5          | 4             | 0.632              | 81.565                       | 73.408     | 0.407    | 36.819               | 36.819                      |  |
| 6          | 8             | 0.482              | 61.750                       | 55.575     | 0.308    | 73.716               | 73.716                      |  |
| 7          | 16            | 0.104              | 11.810                       | 10.629     | 0.059    | 55.634               | 55.634                      |  |
| 8          | 24            | 0.096              | 10.751                       | -1.372     | 0.0056   | 10.634               | 10.634                      |  |
| -          |               |                    |                              | Table No.6 |          |                      |                             |  |
| C N        | Time          | Absorbance         | Concentration                |            |          | Cumulative           | %Cumulativ                  |  |
| S.No       | (Hrs)         | (nm)               | (µg/ml) (X)                  | X*900/1000 | 5*X/1000 | Amount               | drug release                |  |
| 1          | 0             | 0                  | 0                            | 0          | 0        | 0                    | 0                           |  |
| 2          | 0.5           | 0.158              | 18.949                       | 17.054     | 0.094    | 5.208                | 5.208                       |  |
| 3          | 1             | 0.261              | 32.226                       | 29.300     | 0.0162   | 17.216               | 17.216                      |  |
| 4          | 2             | 0.345              | 43.652                       | 39.286     | 0.218    | 29.518               | 29.518                      |  |
| 5          | 4             | 0.497              | 63.731                       | 57.357     | 0.318    | 39.604               | 39.604                      |  |
| 6          | 8             | 0.365              | 46.494                       | 41.664     | 0.231    | 57.588               | 57.588                      |  |
| 7          | 16            | 024                | 30.706                       | 27.635     | 0.153    | 41.817               | 41.817                      |  |
| 8          | 24            | 0.183              | 22.252                       | 20.026     | 0.111    | 27.746               | 27.746                      |  |
| Table No.7 |               |                    |                              |            |          |                      |                             |  |
| a N        | Time          | Absorbance         | Concentration                |            |          | Cumulative           | %Cumulativ                  |  |
| S.No       | (Hrs)         | (nm)               | (µg/ml) (X)                  | X*900/1000 | 5*X/1000 | Amount               | drug release                |  |
| 1          | 0             | 0                  | 0                            | 0          | 0        | 0                    | 0                           |  |
| 2          | 0.5           | 0.135              | 15.911                       | 14.319     | 0.079    | 1.57                 | 1.57                        |  |
| 3          | 1             | 0.21               | 24.630                       | 22.167     | 0.123    | 14.442               | 14.442                      |  |
| 4          | 2             | 0.314              | 39.557                       | 35.6013    | 0.178    | 22.345               | 22.345                      |  |
| 5          | 4             | 0.643              | 83.018                       | 74.716     | 0.373    | 35.974               | 35.974                      |  |
| 6          | 8             | 0.405              | 51.578                       | 46.420     | 0.232    | 74.648               | 74.648                      |  |
| 7          | 16            | 0.302              | 37.972                       | 34.174     | 0.170    | 46.59                | 46.59                       |  |
| 8          | 24            | 0.012              | -0.336                       | -0.302     | -0.0015  | 34.172               | 34.172                      |  |
| 0          | 2.            | 0.012              |                              | Table No.8 | 0.0012   | 51.172               | 01.172                      |  |
| S.No       | Time<br>(Hrs) | Absorbance<br>(nm) | Concentration<br>(µg/ml) (X) | X*900/1000 | 5*X/1000 | Cumulative<br>Amount | %Cumulative<br>drug release |  |
| 1          | 0             | 0                  | 0                            | 0          | 0        | 0                    | 0                           |  |
| 2          | 0.5           | 0.293              | 36.783                       | 33.104     | 0.183    | 2.258                | 2.258                       |  |
| 3          | 1             | 0.328              | 41.406                       | 37.265     | 0.207    | 33.311               | 33.311                      |  |
| 4          | 2             | 0.489              | 62.675                       | 56.407     | 0.313    | 37.578               | 37.578                      |  |
| 5          | 4             | 0.551              | 70.871                       | 63.783     | 0.354    | 56.761               | 56.761                      |  |
| 6          | 8             | 0.464              | 59.372                       | 53.434     | 0.296    | 64.079               | 64.079                      |  |
| 7          | 16            | 0.381              | 48.408                       | 43.567     | 0.242    | 53.676               | 53.676                      |  |
| 7          | 10            |                    |                              |            |          |                      |                             |  |

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|      |            | No.9: %Cumulative | %Cumulative     | %Cumulative     | %Cumulative     | %Cumulative     |
|------|------------|-------------------|-----------------|-----------------|-----------------|-----------------|
|      | <b>T</b> . |                   |                 |                 |                 |                 |
| S.No | Time       | drug release of   | drug release of | drug release of | drug release of | drug release of |
|      |            | <b>F1</b>         | F2              | F3              | F4              | F5              |
| 1    | 0          | 0                 | 0               | 0               | 0               | 0               |
| 2    | 0.5        | 5.433             | 10.414          | 5.208           | 1.57            | 2.258           |
| 3    | 1          | 60.274            | 25.673          | 17.216          | 14.442          | 33.311          |
| 4    | 2          | 20.885            | 32.321          | 29.518          | 22.345          | 37.578          |
| 5    | 4          | 26.614            | 36.819          | 39.604          | 35.974          | 56.761          |
| 6    | 8          | 52.339            | 73.716          | 57.588          | 74.648          | 64.079          |
| 7    | 16         | 38.484            | 55.634          | 41.817          | 46.59           | 53.676          |
| 8    | 24         | 26.979            | 10.634          | 27.746          | 34.172          | 44.037          |

Table No. 9: % Cumulative drug release of all formulations (F1 F2 F3 F4 and F5)

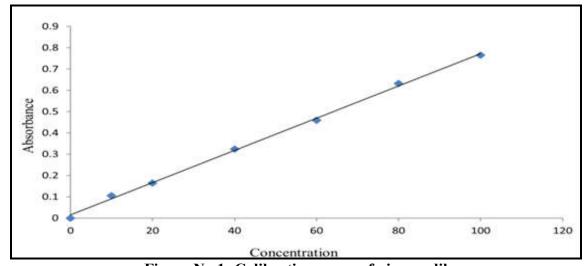
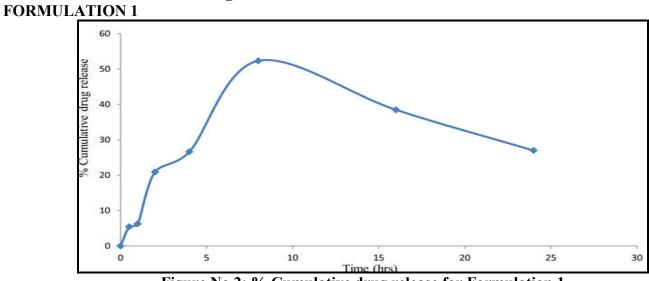
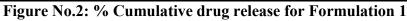


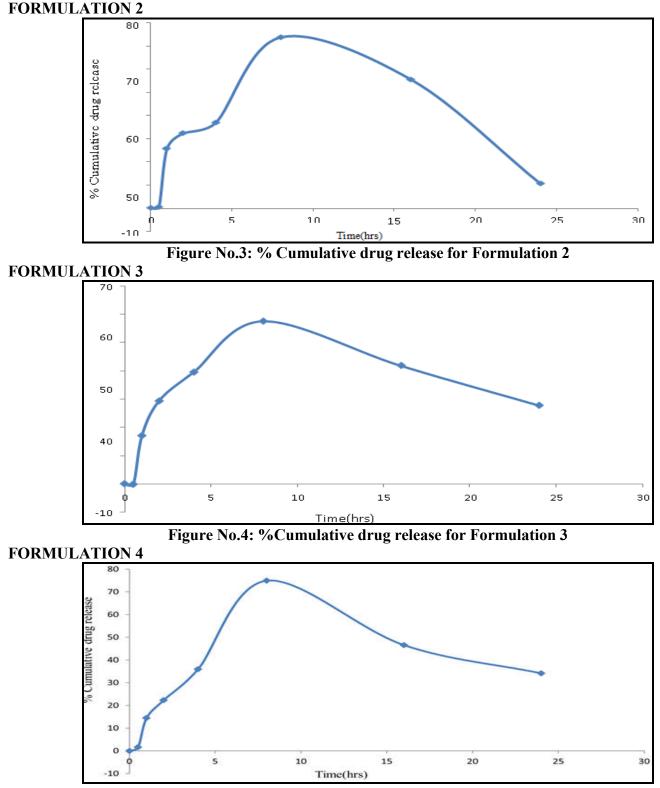
Figure No.1: Calibration curve of nicorandil

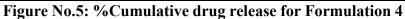




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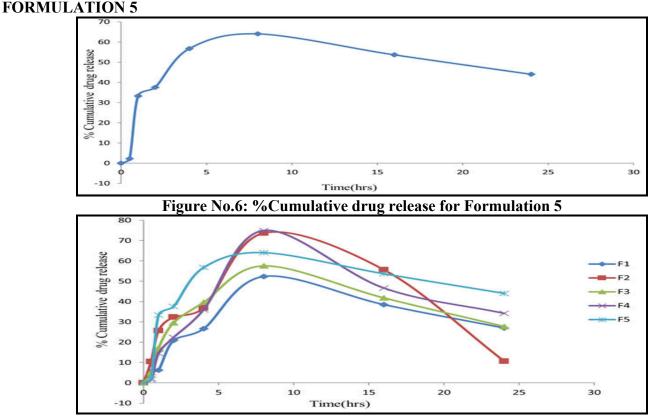


Figure No.7: Cumulative drug release of all formulations (F1, F2, F3, F4, and F5)

#### CONCLUSION

The present investigation revealed that *Tamarindus kernels* mucilage and guar gum, Ethyl cellulose and HPMC appears to be suitable for use as a release retarding agents in the formulations. From the dissolution study, it was concluded that *Tamarindus kernals* mucilage can be used as an excipient for making sustained release matrix tablets of Nicorandil.

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#### **CONFLICT OF INTEREST**

We declare that we have no conflict of interest.

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