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# Research Article

# Development of Metaclopramide Oral Disintegrating Tablets by Using Super Disintegrants

# S. Revathi<sup>\*1</sup>, Saddapalli Baba Fakroddin, Yerradoddy Mamatha, Pachigalla Palvijetha, Sannareddy Pavan Saireddy, Dandu Manisha bee

<sup>1</sup>Associate Professor, Department of Pharmaceutics, Ratnam Institute of Pharmacy, Pidathapolur(V), Muthukur (M), Nellore-524346, SPSR Nellore (Dist) Andhra Pradesh, India.

<sup>2</sup>*Ratnam Institute of Pharmacy, Pidathapolur(V), Muthukur (M), Nellore-524346, SPSR Nellore (Dist) Andhra Pradesh, India.* 

#### Abstract

The mouth dissolving tablet computers (ODTs) actually contains clonidine active ingredient (mcp), had been designed to use variety of different preparations to reinforce it and levels of comfort. Its leakages seem to be merged to group was made up but instead sodium acetate glyceryl there as content after all 3%, 5% and seven percentage w/w. This would be accomplished to choose but also maximise an accumulation of best power splitting criterion getting that whole lesser -ir moments or swiftest solubility. That whole rheological properties of pellet blends have been analyzed employing carr's score, angular position yeah oeuvre and also the hausner's percentage. It and tablet devices seemed to be assessed just that load variability, surface area, compressive strength, occurrence, solid dispersion and now in vitro degradation. It and perseverance of all the most effective mode or optimum value like so segmentations as an oral form imploding tablet computers must have been produced through it compression molding. The ultimate analysis of data discovered that perhaps a pairing like ethyl cellulose, glucose but also 7% group was made up made the highest generic version because once assessor.

**Keywords:** Part is responsible, clonidine active ingredient, sorbitol, mouth dissolving tablet devices or sodium polyethylene glycol.

#### Article Info

Corresponding Author: S. Revathi Associate Professor, Department of Pharmaceutics, Ratnam Institute of Pharmacy, Pidathapolur(V), Muthukur (M), Nellore-524346, SPSR Nellore (Dist), Andhra Pradesh, India.

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#### Contents

| 1. | Introduction            |
|----|-------------------------|
| 2. | Methodology             |
| 3. | Pharmacological studies |
| 4. | Conclusion              |
| 5. | References              |

#### 1. Introduction

Tablet dosage aspects monopolize the most important and also the most prominent position within all dosage forms applications, and so it regarded one of its most popular product delivery methods. Use of a glass of warm watering accept a smart phone is indeed the quickest but also most appropriate method after all government of a narcotic to just a customer. There are many widths akin to its dosage which can then be specified or the contour of something like the laptop. A little laptops might well consist as little as 1000 mg which is less than this same narcotic, others and encompass 100 mg. Of an involved each for a device, that also provides a massive wide range of therapeutic product. Tablet and capsule applications can just be available in different dimensions, and or the opioid excipient could comprise zero<sup>1-5</sup>. 1% of about 90% of either a touch screen mass. It is also very straightforward experience to fabricate devices compared with certain other dosage. Touch screen manufacturing could perhaps achieve the utmost yield each for a fabrication day or and it is the just about all cost effective, specially if someone deems new industrial techniques including process after all compression molding.

#### **Criteria for Orally Disintegrating Tablets**

- Odt's dissolve/disintegrate in few minutes at a time too before positioned there in teeth, and as a direct consequence can indeed be considered but without the demand anyway oceans.
- They are suitable as for edible coatings and many other components.
- Odts continue providing decent consistency till individuals illustrate fewer tolerance to environmental situations.

#### **Pharmacokinetics of ODTs**

Drug will just be assimilated or achieve it and medicinal echelon as well as official launch this same medicinal intervention when it located here on throat. Therefore, its frequency but rather lengthened permeation are indeed very significant. Finished dosage type of smart phone seems to have a lengthier dissolution rate may well ultimately leads complete slow inside this solubilization publication. Through comparision, totally destroyed this same superdisintegrants but rather separation publication are that much quickly. Its solubility may well open here on teeth, but then just laryngeal but also these it till it liquid achieve that whole throat.

#### Various Techniques Used in ODTs Manufacturing

- Freeze Drying/ Lyophilization
- Tablet Moulding
- Spray Drying
- Melt Granulation
- Sublimation
- Direct Compression
- Mass Extrusion
- Taste Masking
- Cotton Candy Process

#### **Direct Compression**

For most drug makers, direct compression technique denotes the best and also most value development of manufacturing technic. They'll be using the traditional machining hardware cos of the helped to improve preparations – particularly mega segmentations but also fructose components the said managed to make this method to be far more trustable with in closed down time to prepare can provide great physical resilience or a faster vitro dissolution.

#### 2. Material and Methods

Metoclopramide was purchased free sample from Piramal health care limited, Microcrystalline cellulose, Mannitol, Crospovidone, sodium starch glycolate, Talc and magnesium stearate was purchased from S.D Fine Chemicals Hyderabad and other ingredients victimized in with analytical grades<sup>6-9</sup>.

#### Preparation and evaluation of metoclopromide ODTs:

In the current investigation, totally destroyed like encountered employing distinction soo segmentations seem to have been willing per the algebraic expressions reported in table six. Two via using compression molding particular method but rather judged just that demonstrate a high degree attributes, production/post tensional metrics and that in vitro release study assets.

#### Evaluation of micromeritic properties of the blends: Angle of repose:

Angle of repose is determined by using funnel method. The accurately weighed blend is taken in a funnel. The height of the funnel is adjusted in such a way that the tip of the funnel just touches the apex of the heap of blend. The drug (as solid dispersion) – excipient blend is allow to flow through the funnel freely on to the surface. The diameter of the powder cone is measured and angle of repose is calculated using the following equation.

#### $Tan \Theta = h/r$

Where h and r are the height of cone (cm) and radius cone base (cm) respectively. Angle of repose less than  $30^{\circ}$  shows the free flowing of the material.

#### **Bulk density:**

Apparent bulk density is determined by pouring a weighed quantity of blend into graduated cylinder and measuring the volume and weight. Bulk density can be calculated by using following formula:

# Bulk density = Weight of the powder / Volume of the packaging

#### **Tapped density:**

It is determined by placing a graduated cylinder, containing a known mass of drug excipients blend. The cylinder is allowed to fall under its own weight onto a hard surface from the height of 10 cm at 2 second intervals. The tapping is continued until no further change in volume is noted. Tapped density can be calculated by using following formula:

# Tapped density = (Weight of the powder / volume of the tapped packing)

#### Carr's index / Compressibility index:

It indicates powder flow properties. It is expressed in percentage and can be calculated by using following formula:

# Compressibility Index (%) = $\frac{TD-BD}{TD}$ x 100

Where, TD is the tapped density of the powder blend and BD is the bulk density of the powder blend.

#### Hausner's ratio:

A similar index to indicate the flow properties can be defined by Hausner's ratio. Hausner's ratio can be calculated by using following formula:

Hausner's ratio = 
$$\frac{\text{Tapped density x 100}}{\text{Poured density}}$$

Hausner's ratio < 1.25 – Good flow = 20% compressibility index

1.25 – Poor flow = 33% compressibility index

#### **Evaluation of Prepared MCP ODTS:**

The compressed ODTs were evaluated for following properties:

#### Uniformity of weight of tablets:

10 tablets were selected from each batch and individually weighed. The average weight and standard deviation of 10 tablets was calculated. The batch passes test for weight variation if not more than two of the individual tablets weight deviate from the average by more than the percentage shown in Table 6.4.

#### Hardness:

The hardness of the MCP ODTs tablets was measured with a Monsanto Hardness Tester (M/s Campbell Electronics, model EIC-66, India). For each batch, 3 tablets were tested. **Friability:** 

For each formulation 10 tablets were weighed, placed in Fryolator (M/S Campbell Electronics, India) and were subjected to 100 rotations in 4 min. The tablets were reweighed, and friability was calculated along with mean and the standard deviation.

#### Friability = W1-W2 / W1 ×100

Where " $W_1$ " is the initial weight and " $W_2$ " is the final weight of the tablets.

#### Drug content:

Ten tablets were weighed and powdered in a mortar. Powder blend equivalent to 10mg of MCP was weighed accurately and transferred to a 10ml volumetric flask, and the MCP was extracted into 10ml methanol. This solution was filtered through  $0.45\mu$  nylon disc filter and the filtered samples were suitably diluted with 1.2 pH buffer and the MCP was assayed by measuring the absorbance at 273nm.

#### Drug – Excipients compatibility studies: FTIR studies

Samples were analyzed using an ATR-FTIR spectrometer (Bruker, Germany). FTIR spectra were measured over the wave number range of 4000-500 cm<sup>-1</sup> at a resolution of 1.0

cm<sup>-1</sup>. The powder sample is placed onto the ATR crystal and the sample spectrum is collected.

# In vitro Disintegration studies:

*In-vitro* disintegration time for tablets was determined using disintegration test apparatus as per USP specifications in 900ml of pH 1.2 buffer. Temperature was maintained at  $37\pm2$  °C. Time for complete disintegration of the tablet with no palpable mass remaining in the apparatus was measured.

#### In vitro Drug release studies:

In-vitro dissolution studies of MCP formulations were carried in 900 mL of pH 1.2 buffer as dissolution medium using USP apparatus II (Paddle method) Dissolution Rate Test Apparatus (DISSO 8000, LABINDIA) with agitation speed of 50 rpm. The temperature was maintained constantly at  $37 \pm 0.5$ °C. 5 mL aliquots were withdrawn at different time intervals and filtered using a 0.45µ nylon disc filters and replaced with 5mL of fresh dissolution medium. The filtered samples were suitably diluted and analyzed at 273nm using UV-Visible Elico SL150 spectrophotometer. Dissolution experiments were conducted in triplicate.

#### First-order release rate constant:

The first order release equation describes the release from systems where the dissolution rate is dependent upon the concentration of the dissolving species. Release behavior generally follows the following first order release equation. A graph of log concentration of drug remaining vs. time yields a straight line with a negative slope.

$$Log C = log C_0 - K_t/2.303$$

Where,

C is the amount of drug undissolved at time t,  $C_0$  is the amount of drug undissolved at t=0,

Kt is the First order release rate constant.

#### 3. Results and Discussion

Analytical method for the analysis of MCP in matrix tablets: UV Spectrophotometric method based on the measurement of absorbance at 273nm was used in the present research work for estimation of MCP. The present analytical method obeyed Beer's law in the concentration range of 2-10µg/mL and is suitable for the estimation of MCP from *in vitro* dissolution samples. The value of  $R^2$  (regression coefficient) for the linear regression equation was found to be 0.997. The linear regression equation for calibration curves is as follows:

#### Y=0.0444x-0.0221

The standard deviation (SD) in the estimated absorbance values was less than 1.5%. These low SD values indicated the reproducibility of the analytical method. The results of standard curve (given in Table 7.1 and shown in Fig 7.1) indicated that the present analytical method was suitable for the estimation of MCP.



Figure 7.1 Calibration curve of MCP in 0.1N HCl

# Evaluation of micromeritic properties of the powder blends:

#### Angle of repose:

Azimuthal after all reduced contact is decided using siphon process. This same weighed and placed mix seems to be considered in some kind of a drip. The peak of a divert was indeed changed in just such a path that nozzle of something like the customer journey only reaches an ascent of both the boatload like integrate. The outcomes got such as table 7.2.

#### Bulk and Tapped densities:

Evident weight ratio is set through it spilling one taken into consideration amount like mix in to one of cylinder or measurement system the quantity as well as pounds. The outcomes got throughout table 7.2

Tapped intensity is set along putting the one measuring pipette, containing volume like painkiller leakages mix. A touching is sustained until it no additional deformation seems to be mentioned.

#### Carr's index and Hausner's ratio:

The powder blends of all the formulations had compressibility index between 18.29 and 21.35 which indicating good flowability of the powder blend. Hausner's ratio for all formulation was less than 1.07, indicating good flowability. The results were given in Table 7.2.

#### **Evaluation of Prepared MCP ODTS:**

#### Uniformity of weight of tablets:

10 tablets were selected from each batch and individually weighed. The average weight and standard deviation of 10 tablets was calculated. The weight of all the tablets was found to be uniform with low standard deviation values indicating efficient mixing of drug, disintegrants and excipients<sup>10-12</sup>.

#### Hardness and Friability:

The hardness was found to be in the range of 3.1 to 3.4 kg/cm<sup>2</sup> for all the formulations indicating good mechanical strength with an ability to withstand physical and mechanical stress conditions while handling. The results were given in Table 7.3. In all the formulations the friability values are less than 1% and meet the IP limits. Friability of the tablets was found below 0.59 - 0.80 % indicating good mechanical resistance of tablets.

**Drug Content:** 

The percentage drug contents of all the tablets were found between 98.23±0.50 percent to 100.23±0.57 percent. The results were given in Table 7.3.

# Drug – Excipients compatibility studies: *FTIR studies*

Samples were analyzed using an ATR-FTIR spectrometer (Bruker, Germany). FTIR spectra were measured over the wave number range of 4000-500 cm<sup>-1</sup> at a resolution of 1.0 cm<sup>-1</sup>. The powder sample is placed onto the ATR crystal and the sample spectrum is collected. The MCP showed characteristic peaks at 1597.06cm<sup>-1</sup> (C=C aromatic stretching), 839.03 cm<sup>-1</sup> (C-C stretching), 3199.91cm<sup>-1</sup> (N-H tertiary amine stretching), 1141.86cm<sup>-1</sup> (aromatic C-O stretching) and 2943.37cm<sup>-1</sup> (C-H aromatic stretching). These characteristic peaks of MCP were all retained in the formulations (F1-F6). These results indicate that there is no interaction between MCP and excipients in formulations. The spectra of MCP and MCP formulations were shown in figures 7.2-7.5.

#### In vitro disintegration studies:

In-vitro disintegration time for tablets was determined using disintegration test apparatus as per USP specifications in 900ml of pH 1.2 buffer. Temperature was maintained at  $37\pm2$  °C. Time for complete disintegration of the tablet with no palpable mass remaining in the apparatus was measured. The results were given in Table 7.5. From the results, it was observed that increase in concentration of the disintegrating agents resulted in faster disintegration of the tablets. Faster disintegration times were observed with formulations containing SSG as disintegrating agent compared to the crospovidone. This might be due to the more swelling ability of the SSG compared to crospovidone.



Figure 7.2 FTIR Spectrum of MCP

#### In vitro disintegration studies:

*In-vitro* disintegration time for tablets was determined using disintegration test apparatus as per USP specifications in 900ml of pH 1.2 buffer. Temperature was maintained at  $37\pm2$  °C. Time for complete disintegration of the tablet with no palpable mass remaining in the apparatus was measured. The results were given in Table 7.5.

From the results, it was observed that increase in concentration of the disintegrating agents resulted in

faster disintegration of the tablets. Faster disintegration times were observed with formulations containing SSG as disintegrating agent compared to the crospovidone. This might be due to the more swelling ability of the SSG compared to crospovidone.

#### In vitro drug release studies:

In the present study the effect of different super disintegrating agents and their concentrations on MCP ODTs was studied. Drug release studies were carried out using 0.1N HCl as dissolution medium. Cumulative percent of MCP released at different time intervals were given in Tables 7.6-7.11 and the drug release profiles were shown in Figs 7.6-7.11.

Initially, formulations F1, F2 and F3 were prepared using crospovidone as disintegrating agent at 3, 5 and 7% w/w levels respectively. The cumulative percent of MCP released was 100.65±0.54, 100.70±1.70 and 100.01±0.39 for F1, F2 and F3 formulations respectively at the end of 720, 600 and 480sec. The results revealed that increase in concentration of crospovidone increase the MCP release rate from the tablets.

Further, formulations F4, F5 and F6 were prepared using sodium starch glycolate (SSG) as disintegrating agent at 3, 5 and 7% w/w levels respectively. The cumulative percent of MCP released was 100.68±0.73, 100.58±1.08 and 100.40±0.19 for F4, F5 and F6 formulations respectively at the end of 600, 420 and 240sec. The results revealed that increase in concentration of SSG also increase the MCP release rate from the tablets.

Among the two disintegrating agents used SSG showed faster MCP release rates compared to crospovidone (Figure 7.14). This may be due to the more swelling ability

of SSG resulting in faster disintegration of tablets and in turn the MCP release rates.

Overall, from the results obtained it can be concluded that the type and concentration of super disintegrating agents had significant impact on the MCP release rates with F6 formulation containing 7% w/w of SSG showed superior MCP release rates compared to other formulations.

#### In vitro release kinetics:

To better understand the release profiles obtained with MCP formulations, the drug release data obtained at different time points was fitted in to kinetic models such as First order (Lapidus H et al; 1966), Higuchi model (Higuchi T 1963).

#### Log Qt unreleased versus t (first order) Qt versus square root of t (Higuchi)

Where Qt is the cumulative % MCP released at time t,

The first order release rate constant 'k' (sec<sup>-1</sup>) values and correlation coefficient ( $R^2$ ) values calculated from dissolution data (0-420 sec) for F1-F3 formulations and from 0-300sec for F4-F6 formulations. The results were given in Table 7.12 and the profiles were shown in Fig 7.15 to 7.20.

The 'k' values were higher for F3 formulation compared to F1, F2 formulations. A 2.3 & 4.1 fold increase in 'k' values was observed for F3 compared to that of F1 & F2. Among F4, F5 and F6 formulations the 'k' values were higher for F6 formulation compared to F4 and F5 formulations.

The Higuchi square root model of all MDFs showed higher correlation coefficient values (0.844-0.978) indicating diffusion as release mechanism. The results are given in Table 7.13 and profiles were shown in Fig 7.21 to 7.26.

| Ingredients (mg/tablet) | F1  | F2  | F3  | F4  | F5  | F6  |
|-------------------------|-----|-----|-----|-----|-----|-----|
| MCP HCI                 | 10  | 10  | 10  | 10  | 10  | 10  |
| MCC                     | 40  | 40  | 40  | 40  | 40  | 40  |
| Mannitol                | 43  | 41  | 39  | 43  | 41  | 39  |
| Crospovidone            | 3   | 5   | 7   | -   | -   | -   |
| SSG                     | -   | -   | -   | 3   | 5   | 7   |
| Talc                    | 3   | 3   | 3   | 3   | 3   | 3   |
| Magnesium stearate      | 1   | 1   | 1   | 1   | 1   | 1   |
| Total                   | 100 | 100 | 100 | 100 | 100 | 100 |

#### Table 1: Formulae of the MCP ODTs with different super disintegrating agents

| S.No | Flow properties | Angle of repose (Θ)(degrees) | Compressibility index (%) | Hausner's ratio |
|------|-----------------|------------------------------|---------------------------|-----------------|
| 1    | Excellent       | 25-30                        | <10                       | 1.00-1.11       |
| 2    | Good            | 31-35                        | 1115                      | 1.12-1.18       |
| 3    | Fair            | 34-40                        | 16-20                     | 1.19-1.25       |
| 4    | Possible        | 41-45                        | 21-25                     | 1.26-1.34       |
| 5    | Poor            | 45-46                        | 26-31                     | 1.35-1.45       |

S. Revathi, et al. W. J. Pharm. Biotech., 2023, 10(1): 08–17

| 6 | Very poor | 55-56 | 32-37 | 1.46-1.59 |
|---|-----------|-------|-------|-----------|
| 7 | Very poor | >66   | >38   | >1.6      |

#### Table 3: Weight variation allowed as USPXX-NF XV

| Average weight of tablet (mg) | Percentage difference allowed |
|-------------------------------|-------------------------------|
| ≤130                          | 10                            |
| 130-324                       | 7.5                           |
| >324                          | 5                             |

### Table 7.1 Calibration curve of MCP in 0.1N HCl (n=3)

| S.no Concentration (µg/mL)              |                | Absorbance   |  |  |
|---|----------------|--------------|--|--|
| 1.                                      | 2              | 0.115±0.051  |  |  |
| 2.                                      | 2. 4 0.200±0.0 |              |  |  |
| 3.                                      | 6              | 0.279±0.004  |  |  |
| 4.                                      | 8              | 0.3898±0.001 |  |  |
| 5.                                      | 10             | 0.461±0.032  |  |  |
| Y=0.0444x-0.0221, R <sup>2</sup> =0.997 |                |              |  |  |

# Table 4: Micromeritic properties of the powder blends

| Powder<br>Blend | Bulk density<br>(ρ <sub>ь</sub> ) | Tapped density<br>(ρ <sub>t</sub> ) | Angle of<br>Repose (Θ) | Compressibility<br>index (%) | Hausner's<br>ratio |
|-----------------|-----------------------------------|-------------------------------------|------------------------|------------------------------|--------------------|
| F1              | 0.364±0.01                        | 0.515±0.03                          | 25.34±1.22             | 20.24±0.24                   | 1.30±0.02          |
| F2              | 0.373±0.02                        | 0.534±0.02                          | 23.22±0.99             | 19.75±0.19                   | 1.07±0.01          |
| F3              | 0.378±0.02                        | 0.524±0.01                          | 25.19±1.22             | 15.17±0.18                   | 1.12±0.03          |
| F4              | 0.369±0.03                        | 0.506±0.01                          | 28.75±0.99             | 18.29±0.33                   | 1.16±0.01          |
| F5              | 0.399±0.04                        | 0.590±0.03                          | 29.24±0.89             | 20.75±0.24                   | 1.15±0.02          |
| F6              | 0.411±0.042                       | 0.601±0.02                          | 22.46±0.13             | 21.35±0.34                   | 1.17±0.01          |

# Table 5: Physicochemical properties of the prepared MCP ODTs

| Formulation | Hardness<br>(kg/cm <sup>2</sup> ) | Friability<br>(%) | Weight variation<br>(mg) | Drug content<br>(%) |
|-------------|-----------------------------------|-------------------|--------------------------|---------------------|
| F1          | 3.2±0.26                          | 0.59              | 200±1.15                 | 100.23±0.57         |
| F2          | 3.4±0.43                          | 0.77              | 199±1.17                 | 98.14±0.75          |
| F3          | 3.0±0.21                          | 0.80              | 200±1.05                 | 99.56±0.23          |
| F4          | 3.1±0.28                          | 0.66              | 200±0.54                 | 99.56±0.23          |
| F5          | 3.4±0.43                          | 0.60              | 200±1.08                 | 98.23±0.50          |
| F6          | 3.4±0.71                          | 0.63              | 200±1.03                 | 98.26±0.72          |

## Table 6: FTIR Spectral data of MCP

| Functional group | Vibration  | Frequency range | Frequency Obtained for<br>MCP (cm <sup>-1</sup> ) |
|------------------|------------|-----------------|---|
| C-C              | Stretching | 700-1300        | 837.11  |
| N-H              | Stretching | 3200-3500       | 3263.14   |
| C-0              | Stretching | 1000-1300       | 1147.37   |
| C-H              | Stretching | 2850-3000       | 2918.22   |

### Table 7: Mean dissolution time of MCP tablets (n=3)

| Formulation | Mean Disintegration time (Sec)±SD |
|-------------|-----------------------------------|
| F1          | 189±0.96                          |
| F2          | 108±1.25                          |
| F3          | 54±1.98                           |
| F4          | 130±0.88                          |
| F5          | 86±0.59                           |
| F6          | 43±1.96                           |

# S. Revathi, et al. W. J. Pharm. Biotech., 2023, 10(1): 08–17

|            | Cum     | Mean ± SD |         |             |
|------------|---------|-----------|---------|-------------|
| Time (sec) | Trial-1 | Trial-2   | Trial-3 |             |
| 0          | 0.00    | 0.00      | 0.00    | 0.00±0.00   |
| 15         | 5.26    | 5.11      | 5.96    | 5.44±0.370  |
| 30         | 12.68   | 12.99     | 12.37   | 12.68±02.53 |
| 60         | 19.57   | 18.99     | 19.65   | 19.40±0.29  |
| 90         | 26.48   | 26.85     | 27.02   | 26.78±0.22  |
| 120        | 35.78   | 35.88     | 35.02   | 35.56±0.38  |
| 180        | 41.32   | 41.03     | 41.86   | 41.40±0.34  |
| 240        | 48.7    | 48.46     | 48.99   | 48.71±0.21  |
| 300        | 62.13   | 62.44     | 62.8    | 62.45±0.27  |
| 360        | 69.58   | 69.14     | 69.92   | 69.54±0.31  |
| 420        | 75.9    | 75.17     | 75.5    | 75.52±0.29  |
| 480        | 81.6    | 81.3      | 81.9    | 81.6±0.24   |
| 540        | 88.54   | 88.2      | 88.81   | 88.51±0.24  |
| 600        | 93.26   | 93.56     | 93.88   | 93.56±0.25  |
| 660        | 97.5    | 96.85     | 96.2    | 96.8±0.53   |
| 720        | 100.36  | 100.65    | 100.96  | 100.65±0.54 |

## Table 8: Cumulative percent of MCP released from F1 formulation (n=3)

# Table 9: Cumulative percent of MCP released from F2 formulation (n=3)

|            | Cumulative % of MCP released |         |         |             |
|------------|------------------------------|---------|---------|-------------|
| Time (sec) | Trial-1                      | Trial-2 | Trial-3 | Mean ± SD   |
| 0          | 0.00                         | 0.00    | 0.00    | 0.00±0.00   |
| 15         | 8.79                         | 7.99    | 13.36   | 10.05±2.90  |
| 30         | 12.81                        | 12.02   | 12.5    | 12.44±0.40  |
| 60         | 20.76                        | 20.12   | 21.9    | 20.93±0.90  |
| 90         | 26.98                        | 26.12   | 26.75   | 26.62±0.45  |
| 120        | 37.56                        | 37.2    | 37.8    | 37.52±0.30  |
| 180        | 48.71                        | 48.52   | 48.95   | 48.73±0.22  |
| 240        | 57.36                        | 57.11   | 57.66   | 57.38±0.28  |
| 300        | 63.41                        | 64.01   | 63.56   | 63.66±0.31  |
| 360        | 71.59                        | 71.65   | 72.2    | 71.81±0.34  |
| 420        | 79.98                        | 79.2    | 79.5    | 79.56±0.39  |
| 480        | 85.14                        | 85.5    | 85.9    | 85.51±0.38  |
| 540        | 91.58                        | 92.17   | 95.67   | 93.14±2.21  |
| 600        | 99.26                        | 100.26  | 102.58  | 100.70±1.70 |

# Table 10: Cumulative percent of MCP released from F3 formulation (n=3)

| Time  | Cumulative % of MCP released |         |         | Mean ± SD   |
|-------|------------------------------|---------|---------|-------------|
| (sec) | Trial-1                      | Trial-2 | Trial-3 |             |
| 0     | 0.00                         | 0.00    | 0.00    | 0.00±0.00   |
| 15    | 11.23                        | 12.56   | 12.56   | 12.12±0.63  |
| 30    | 20.57                        | 20.12   | 20.12   | 20.27±0.21  |
| 60    | 31.59                        | 31.69   | 31.69   | 31.66±0.05  |
| 90    | 45.28                        | 45.87   | 45.87   | 45.67±0.28  |
| 120   | 59.12                        | 58.74   | 58.74   | 58.87±0.18  |
| 180   | 67.51                        | 67.87   | 67.87   | 67.75±0.17  |
| 240   | 74.56                        | 74.69   | 74.69   | 74.65±0.06  |
| 300   | 80.54                        | 80.1    | 80.1    | 80.25±0.21  |
| 360   | 89.74                        | 89.99   | 89.99   | 89.91±0.12  |
| 420   | 95.67                        | 95.39   | 95.39   | 95.48±0.13  |
| 480   | 100.56                       | 99.74   | 99.74   | 100.01±0.39 |

# S. Revathi, et al. W. J. Pharm. Biotech., 2023, 10(1): 08–17

|            | Cumulative % of MCP released |         |         |             |
|------------|------------------------------|---------|---------|-------------|
| Time (sec) | Trial-1                      | Trial-2 | Trial-3 | Mean ± SD   |
| 0          | 0.00                         | 0.00    | 0.00    | 0.00±0.00   |
| 15         | 8.54                         | 7.56    | 8.74    | 8.2±0.56    |
| 30         | 15.24                        | 14.28   | 14.26   | 14.59±0.69  |
| 60         | 21.59                        | 22.36   | 20.98   | 21.64±4.90  |
| 90         | 30.28                        | 38.14   | 29.14   | 32.52±2.95  |
| 120        | 36.87                        | 41.28   | 35.67   | 37.94±4.16  |
| 180        | 44.89                        | 49.56   | 41.27   | 45.24±2.95  |
| 240        | 56.27                        | 57.18   | 52.31   | 55.25±4.16  |
| 280        | 65.78                        | 62.34   | 67.18   | 65.10±2.59  |
| 300        | 72.17                        | 74.59   | 74.15   | 73.64±2.49  |
| 360        | 79.84                        | 81.69   | 82.56   | 81.36±1.29  |
| 420        | 83.64                        | 85.63   | 87.65   | 85.64±1.39  |
| 480        | 89.85                        | 91.27   | 90.01   | 90.38±2.01  |
| 520        | 94.78                        | 96.21   | 95.21   | 95.40±0.78  |
| 600        | 99.58                        | 102.24  | 100.23  | 100.68±0.73 |

# Table 11: Cumulative percent of MCP released from F4 formulation (n=3)

 Table 12: Cumulative percent of MCP released from F5 formulation (n=3)

 Cumulative % of MCP released

|            | Cumulative % of MCP released |         |         |             |
|------------|------------------------------|---------|---------|-------------|
| Time (sec) | Trial-1                      | Trial-2 | Trial-3 | Mean ± SD   |
| 0          | 0.00                         | 0.00    | 0.00    | 0.00±0.00   |
| 15         | 14.25                        | 16.28   | 15.28   | 15.27±1.02  |
| 30         | 24.25                        | 27.59   | 26.17   | 26.00±1.68  |
| 60         | 36.74                        | 34.16   | 35.17   | 35.36±1.30  |
| 90         | 49.58                        | 51.49   | 50.29   | 50.45±0.97  |
| 120        | 56.27                        | 57.21   | 55.39   | 56.29±0.91  |
| 180        | 69.21                        | 71.29   | 70.27   | 70.26±1.04  |
| 240        | 77.89                        | 82.59   | 79.21   | 79.90±2.42  |
| 300        | 87.98                        | 91.28   | 88.27   | 89.18±1.83  |
| 360        | 94.26                        | 95.68   | 95.32   | 95.09±0.74  |
| 420        | 100.59                       | 99.5    | 101.65  | 100.58±1.08 |

#### Table 13: Cumulative percent of MCP released from F6 formulation (n=3)

|            | Cumulative % of MCP released |         |         |             |
|------------|------------------------------|---------|---------|-------------|
| Time (sec) | Trial-1                      | Trial-2 | Trial-3 | Mean ± SD   |
| 0          | 0.00                         | 0.00    | 0.00    | 0.00±0.00   |
| 15         | 16.6                         | 17.89   | 17.10   | 17.19±0.74  |
| 30         | 32.98                        | 34.56   | 32.58   | 33.32±0.9   |
| 60         | 55.65                        | 56.97   | 58.17   | 56.93±0.76  |
| 90         | 72.95                        | 79.1    | 82.60   | 78.21±3.55  |
| 120        | 89.66                        | 83.6    | 84.10   | 85.78±3.50  |
| 180        | 95.82                        | 94.8    | 94.12   | 94.91±0.59  |
| 240        | 100.95                       | 101.28  | 99.10   | 100.40±0.19 |

# Table 14: $DP_{60}$ and First order kinetic data for MCP formulations

| FORMULATION | DP <sub>60</sub> ★<br>(Mean ± SD) | R <sup>2</sup> (First order plot) | Mean 'k' (sec <sup>-1</sup> )<br>(0-420 sec) |
|-------------|-----------------------------------|-----------------------------------|--|
| F1          | 19.14 ± 0.29                      | 0.990                             | 0.0034                                       |
| F2          | 20.92 ± 0.90                      | 0.994                             | 0.0032                                       |
| F3          | 30.83 ± 1.14                      | 0.968                             | 0.0034                                       |
| F4          | 21.64 ± 0.69                      | 0.968                             | 0.0064                                       |
| F5          | 35.36 ± 1.30                      | 0.991                             | 0.0046                                       |

S. Revathi, et al. W. J. Pharm. Biotech., 2023, 10(1): 08-17

| F6 | 56.93 ± 1.26 | 0.997 | 0.0069 |
|----|--------------|-------|--------|
|    |              |       |        |

**DP**<sub>5</sub>★: Drug percent released at 60 sec.

| FORMULATION | R <sup>2</sup> | K <sub>H</sub> (sec <sup>-1/2</sup> ) |
|-------------|----------------|---------------------------------------|
| F1          | 0.979          | 3.891                                 |
| F2          | 0.984          | 4.070                                 |
| F3          | 0.985          | 4.925                                 |
| F4          | 0.971          | 4.169                                 |
| F5          | 0.995          | 5.331                                 |
| F6          | 0.942          | 7.391                                 |

# Table 15: Higuchi plot-R<sup>2</sup> values for MCP formulations

#### Table 16: Drug release kinetics of metoclopramide oral disintegrating tablets

| Batch | FIRST-ORDER<br>R <sup>2</sup> | HIGUCHI<br>R <sup>2</sup> | Release Mechanism |
|-------|-------------------------------|---------------------------|-------------------|
| F1    | 0.9903                        | 0.9799                    | Non-fickian       |
| F2    | 0.9943                        | 0.9849                    | Non-fickian       |
| F3    | 0.9688                        | 0.9859                    | Non-fickian       |
| F4    | 0.9681                        | 0.9711                    | Non-fickian       |
| F5    | 0.9919                        | 0.9953                    | Non-fickian       |
| F6    | 0.997                         | 0.9733                    | Non-fickian       |

#### 4. Conclusion

Metoclopramide active ingredient (mcp) is really a drug always used as a tummy or oesophagus difficulties. It's also commonly prescribed and for nausea, to assist to discharge of something like the belly along individuals with deferred throat discharge, hemolyticanemia as well as to assist as both acid reflux trader gastroesophageal reflux. The drugexcipient suitability must've been revealed whilst also FT-IR research. A conclusions drawn through ft-ir study showed there was no pesticide interplay among opioid but rather components<sup>13</sup>.

Pre-compression parametric for maximum dry density, utilized surface area, stress - strain benchmark, angular velocity sure optimum tilt but also hausner's assets ratio seemed to be estimated for 3 techniques or described in table seven. Two. Its excellent flow nanoparticles were determined to really be decent per the rules limiting. It and tightly compacted capsules had been reviewed rather than different quality testing somewhere around mass fluctuation, stiffness, fracture toughness, pharmacopoeial. Together all regimens had been great as well as cooperates the boundaries who were offered throughout tabular form six. In-vitro super disintegrants just that devices was firm utilizing dissolving monitoring device according the glock specs along 900ml yeah flow rate operand. Two cushion. Through the outcomes, this became witnessed a certain increase in levels of falling to pieces agencies contributed out way quicker dispersion of something like the tablet devices<sup>14-17</sup>. Quickly complete destruction twice seemed to be witnessed to assist in making senior program just like loss of enthusiasm advisor in comparison to this same ratio is the relationship.

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#### **Conflicts of interest**

The authors attest that they have no conflict of interest in this study.

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#### 5. References

- [1] Anilkumar J. Shinde, Amit L. Shinde, Sachin A. Kandekar, Ravindra J. Jarag, Harinath N. More; Formulation and evaluation of fast dissolving tablet of cephalexin using super disintegrant blends and subliming material: Journal of Pharmacy Research: 2009, vol. 2: 1509-1511.
- [2] Ashok Kumar, A. G. Agrawal; Formulation development and evaluation of orally disintegrating tablets by sublimation technique: International Journal of PharmTech Research: 2009, vol. 1: 997-999.
- [3] Avani R. Gosai, Sanjay B. Patil and Krutika K. Sawant; Formulation and evaluation of orodispersible tablets of Ondansetron Hydrochloride by direct compression using superdisintegrants: International journal of pharmaceutical sciences and nanotechnology: 2008, vol. 1: 107-111.
- [4] B.G. Shiyani, R.B. Dholakiya, B.V. Akbari, D.J. Lodhiya, G.K. Ramani; Development and evaluation of novel

immediate release tablets of metoclopramide HCl by direct compression using treated gellan gum as a disintegration-accelerating agent: Journal of Pharmacy Research 2009, vol. 2 (9): 1460-1464.

- [5] Basawaraj S. Patil, K. Dayakar Rao, Upendra Kulkarni; Formulation and Development of Granisetron Hydrochloride Fast Dissolving Tablets by Sublimation Technique: International journal of pharmacy and pharmaceutical science research: 2011, vol. 1: 20-25.
- [6] Bhowmik, D., et al., Fast dissolving tablet: An overview. Journal of chemical and Pharmaceutical Research, 2009. 1(1): p. 163-177.
- [7] Bhunia Biswajit, Varun joshi; Formulation and evaluation of orodispersible tablet of Amlodipine Besylate: International journal of pharmacy and technology 2011, vol. 3: 37453766.
- [8] Bhupendra G. Prajapati, Dipesh V. Patel; Comparative study of efficiency of different superdisintegrant for fast dissolve tablet of domperidone: Journal of Pharmacy Research: 2010, vol. 3: 151-155.
- [9] Chang, R.-K., et al., Fast-dissolving tablets. Pharmaceutical technology, 2000. 24(6): p. 52-58.
- [10] Chue, P., R. Welch, and C. Binder, Acceptability and disintegration rates of orally disintegrating risperidone tablets in patients with schizophrenia or schizoaffective disorder. Canadian journal of psychiatry, 2004. 49(10): p. 701-703.
- [11] Dixit, S., et al., Fast Dissolving Tablet-A Promising Approach for Drug Delivery: A Review. Journal of Pharmacy Research, 2012. 5(3): p. 1508-1513.
- [12] Fass, R., H. Pieniaszek, and J. Thompson, Pharmacokinetic comparison of orally- disintegrating metoclopramide with conventional metoclopramide tablet formulation in healthy volunteers. Alimentary pharmacology & therapeutics, 2009. 30(3): p. 301-306.
- [13] Fu, Y., et al., Orally fast disintegrating tablets: developments, technologies, taste-masking and clinical studies. Critical Reviews<sup>™</sup> in Therapeutic Drug Carrier Systems, 2004. 21(6).
- [14] Gauri, S. and G. Kumar, Fast dissolving drug delivery and its technologies. The pharma innovation, 2012. 1(2): p. 34-39.
- [15] Ghosh, T., A. Ghosh, and D. Prasad, A Review on New Generation Orodispersible Tablets and Its Future Prospective. International Journal of Pharmacy and Pharmaceutical Sciences, 2011. 3(1): p. 1.
- [16] Glare, P., et al., Systematic review of the efficacy of antiemetics in the treatment of nausea in patients with far-advanced cancer. Supportive Care in Cancer, 2004. 12(6): p. 432-440.
- [17] Kumar B. Sutradhar, Dewan T. Akhter, Riaz Uddin; Formulation and evaluation of taste masked oral dispersible tablets of Domperidone using sublimation method: International journal of pharmacy and pharmaceutical sciences 2012, vol. 4: 727-732.