

**Journal of Advanced Pharmaceutical Sciences and Natural Products (JAPSNP)****ORAL DRUG DELIVERY SYSTEMS WITH EMPHASIS ON EXTENDED AND SUSTAINED RELEASE MECHANISMS: A REVIEW****Shivani Sharma**

Sanskar College of Pharmacy  
& Research, Ghaziabad,  
201302, Uttar Pradesh, India

**Anuradha Verma**

Sanskar College of Pharmacy  
& Research, Ghaziabad,  
201302, Uttar Pradesh, India

**Babita Kumar**

Sanskar College of Pharmacy  
& Research, Ghaziabad,  
201302, Uttar Pradesh, India

**Corresponding author:**

Shivani Sharma

**Email:** [shivani22-  
mpharm@sanskar.org](mailto:shivani22-mpharm@sanskar.org)

**ABSTRACT**

Oral drug delivery is one of the most common and convenient ways to give medicine. It involves taking drugs by mouth so they can work either locally in the mouth or throughout the body. One important type of oral delivery is extended release, where the drug is released slowly over time. This method helps keep a steady level of medicine in the body, improving treatment and reducing how often patients need to take the drug. This review discusses different oral drug delivery systems, focusing on extended and sustained release formulations. It explains how these systems work to release drugs gradually, either by controlling the way the drug dissolves or by using special materials that slow down drug release. The mechanisms behind sustained release include diffusion, erosion, and osmotic pressure, among others. Understanding these extended release oral systems is important for improving patient compliance and therapy effectiveness. Advances in materials and technology continue to enhance these drug delivery methods, providing better options for treating chronic diseases and improving patient quality of life.

**Keywords:** Oral Drug Delivery System, Sustained release, Dosage form, Gastrointestinal tract, Extended Release



## Journal of Advanced Pharmaceutical Sciences and Natural Products

### 1. Introduction

Oral conveyance medications by a wide margin the most reasonable medicine delivery process due to its ease of administration, patient consistency, plan flexibility, and other factors. Oral medication delivery systems, including oral medicine administration systems, make for a significant portion of the medication conveyance frameworks available on the market advanced over time from timely delivery to site-specific delivery. Every patient may have a continuing desire for best possible drug delivery system that addresses two basic characteristics of single-portion and less continuous dosing over the course of treatment. The measurements structure should deliver dynamic medication straightforwardly at site of activity. Hence, target of drug specialist is to foster a solitary portion treatment that can be ideal framework as could be expected. Endeavors to foster a solitary portion treatment for entire length treatment stand out on controlled or Broadened Delivery drug conveyance frameworks. As of late in field of drug innovation research have been completed by utilizing existing particle to defeat troubles, for example, unfortunate dissolvability, unfortunate bioavailability, dosing issue, steadiness, poisonousness & so on. This pattern of working lead to the improvement of new medication conveyance framework. Oral medication conveyance framework ought to be essentially pointed toward accomplishing the expanded bioavailability of medications, as the medications bioavailability is a significant calculate

remedial viability of the medication handily ingested (GIT), brief quickly bloodstream, necessitating continuous dosage. In order to avoid this, definitions of oral supported controlled discharge have been created trying to deliver the medication gradually into GIT and keep a viable medication fixation. The important objective of oral controlled discharge conveyance frameworks is to convey medication on time span that will increment adequacy and limit unfriendly impacts.<sup>1-3</sup>

Regular measurement structure, for example, tablets, case produce explicit medication focus without offering any command over drug conveyance framework in contrast with control discharge framework. Different sorts of medication conveyance frameworks for oral organization, for example, rate controlled convey frameworks, time-controlled conveyance frameworks & site-explicit conveyance framework have been widely evolved instances rate controlled & time controlled conveyance framework, supported retention restricted travel season measurements structure assimilation the ingestion site on grounds that, from there on ,the delivered drug isn't retained. Consequently, when a medication has a tight 'Retention window plan of the Drawn out Delivery readiness measurement structure & controlled drug discharge. Drug retention from the gastrointestinal lot is an exceptionally intricate technique and includes m any factors as we as a whole know the degree of ingestion from GIT relies upon contact time with the small digestive system mucosa so the digestive



## Journal of Advanced Pharmaceutical Sciences and Natural Products

system travel is a significant component for drugs that are not completely consumed.<sup>4</sup>

### 2. Gastrointestinal Tract

GIT, often known as human gastrointestinal tract, is an organ system responsible for devouring & processing staples, engrossing supplements, & ousting waste. The parcel comprises of stomach & digestive organs, is parted into plots of upper & lower digestive tracts. Every graphic across mouth butt is incorporated into GI lot. Then again, the stomach related framework is a more extensive term that involves different designs, including stomach related organs.<sup>5,6</sup> The GI lot discharges chemicals to assist with direct-ing stomach related procedure. These substances include ghrelin, cholecystokinin, secretin, and gastrin. The property is divided into upper and lower plots, as well as small and large sections of the digestive tract:

#### 2.1 Upper Gastrointestinal Tract

The upper gastrointestinal system is made up of neck, stomach, & duodenum. The exact separation b/w upper & lower halves of duodenum is sus-pensory tendon, sometimes referred to as Tendon of Treitz. Clinicians frequently use this divide to represent gastrointestinal drainage as having a "upper" or "lower" beginning, and it also delineates the early stage borders across the foregut and midgut.<sup>7,8</sup>

#### 2.2 Lower GIT

The entire internal organ and most of tiny digestive tract are included in the lower gastrointestinal parcel. The digestive system, also known as gut, stomach portion that extends stomach's butt. In humans & other warm-blooded animals, it is divided

into two parts: small digestive system & subsequent digestive organ. In humans, digestive organ is divided into cecum, colon, rectum, & butt-centric channel, while small digestive system is almost certainly divided into duodenum, jejunum & ileum.<sup>9</sup>

### 3. Conventional dosage form

Frequent measurements cause a wide range of changes in way drugs are fixed in tissues and circulatory system, which leads to unintended injury & regrettable productivity. For instance, this component contributed to concept of restricted drug conveyance frameworks due to redundant dosing & flighty retention. Through limitation at place of activity, lowering the portion required, or providing uniform medication conveyance, planning controlled conveyance frameworks medication proficiency. The main goals stretched-out delivery delivery are to ensure safety, advance medicine adequacy, and promote silent consistency. By creating a Broadened Delivery pill, this ought to be feasible.<sup>10,11</sup>

#### Limitations of Conventional Oral Dosage Form

- Seesaw changes; unfortunate patient consistency;
- Likelihood of missing chunk.
- Various medication treatment improves the gamble of harmfulness generally treatment.
- Approaches to overcome these limitations
- Creation of new, safer, & better medications with longer half-lives & a wealth of helpful data;



## Journal of Advanced Pharmaceutical Sciences and Natural Products

- The safe and effective use of medications that are already on the market by applying the ideas and methods of regulated and authorized drug delivery systems.<sup>12</sup>

### 4. Extended Release Drug Delivery System

Broadened discharge plan a significant program for new medication innovative work to meet a few neglected clinical requirements viz. gives increment bioavailability of medication item, decrease in the recurrence of organization to drag out span of compelling blood levels, Diminishes the variance of pinnacle box focus and aftereffects and potentially works on the particular appropriation of the medication.<sup>13,14</sup> Expanded discharge drug conveyance framework accomplishes a sluggish arrival of the medication over a lengthy timeframe or the medication is retained over a more drawn out timeframe. How much medication is delivered at a controlled rate (support portion, DM) to keep up with the specific blood levels for want timeframe.<sup>15</sup>

### Desired Criteria for Extended ERDDS<sup>16,17</sup>

The medications must formed an ERDDS ought to meet following boundaries.

- It should be stable in GIT medium & be able to be taken orally.
- Ideally, a medicine with a half-life of 2-4 hours would be a good candidate to be included in emergency room dosage formulations, such as captopril or salbutamol sulfate.
- Since oral course is suitable for medications given in portions high 1g,

such as metronidazole, portion of medication should be less than 0.5g.

### Merits of Extended ERDDS<sup>18-20</sup>

- The lengthy delivery details might keep up with remedial fixations over delayed periods.
- The utilization of broadened discharge details dodges the high blood fixation.
- Diminish the harmfulness by easing back drug assimilation.
- Limit the neighborhood and foundational secondary effects.
- Further develop treatment viability.
- Limit drug amassing with persistent dosing.
- Upgrade of movement span for short half-life drugs.

### Advantages of Extended Release Drug Delivery<sup>21</sup>

- Further developed treatment.
- Supported blood level.
- Constriction of unfavorable impacts.
- Patient Accommodation/worked on tolerant consistence.
- Economy.
- Expanded Delivery details are more affordable than customary measurements structures.
- Economy may likewise be impacted because of diminished cost of nursing time for organization of medication.
- Blood level swaying normal for multiple dosing of conventional dose structures is decreased.
- Managed portion is diminished.
- Greatest medication accessibility with a base portion.



## Journal of Advanced Pharmaceutical Sciences and Natural Products

- Security edge of high power medications can be expanded.

### Disadvantages<sup>22,23</sup>

- Portion Unloading.
- Less adaptability in intense portion change.
- Poor in vitro - in vivo connection.
- Patient variety.
- Broadened Delivery dose structures are costly.
- It is not recommended to use a medicine with expanded delivery in patients who are known to have impaired kidney or gastrointestinal function.
- It is not appropriate to display medications with extended natural half-lives in an expanded delivery

### 4.1 Types of Extended-release Devices

**4.1.1 Controlled Release (CR):** Controlled discharge frameworks give drug discharge in a sum adequate to keep up with the restorative fixation overstretched time frame.

**Prolong Action:** Draw out or long activity items are measurements structures containing prodrug of remedial substance having delayed natural half-life.

**Sustained Release:** In event of Expanded Delivery (SR) dose shapes the arrival of the medication is more slow than regular measurements structure.<sup>24</sup>

**4.1.2 Delayed Release:** A deferred discharge measurement structure discharges drug at a time other than following organization.<sup>25</sup>

### Theory of Sustained Release

Extended Release Dosage Form Contains:

The loading dose (a) and the maintenance dose (b).

While the maintenance dose, also known as the slowly accessible fraction, releases the medication gradually & sustains therapeutic level for a longer amount of time, loading dose, also known as immediately available portion, quickly reaches therapeutic level after delivery.<sup>28</sup>

### 5. Mechanism of Drug Release from a Sustained Dosage Form

#### Leaching (Diffusion) Type:

A polymeric matrix that is insoluble in water contains the medication. The diffusion is driven by the drug's water solubility within the matrix.

#### Erosion (Dissolution) Type:

Partially water soluble polymers mixture of soluble & in-soluble polymers constitutes matrix. The matrix eroded at various places form which drug will be slowly released.

#### Types of Sustain-release Product:

##### Dissolution-controlled products:

In this approach, drug's rate of dissolution (thus, its availability for absorption) is controlled via slowly soluble polymers & micro-encapsulation process. When the soluble coating & polymer comes into contact with GI fluid, it gradually dissolves, allowing drug to be absorbed. By changing coat's thickness & composition, rate of drug release can be controlled.

#### Erosion Products



## Journal of Advanced Pharmaceutical Sciences and Natural Products

The rate at which the polymer utilized in these items erodes controls drug's release. An example of this formulation is an osmotic pump system, such as Sinemet CR. The steady flow. The rate at which medicine is released is determined by flow of H<sub>2</sub>O through a semi-per-meable membrane into a reservoir that contains an osmotic agent. The medication is either mixed with drug & placed in a reservoir. Osmotic pressure causes dissolved medicine to be pushed from dosage form's opening at a rate of H<sub>2</sub>O ingress.

### Ion Exchange Resins:

Ion exchange resins are used to bind medications. The ionic environment in the gastrointestinal tract controls the drug's release after administration.<sup>29</sup>

### Matrix System:

A solid medication distributed throughout system in this approach. Both the dissolving and diffusion methods regulate the drug's release. The matrix system is the most often utilized technique among the different approaches used to regulate drug release from pharmaceutical dosage forms. The characteristics listed below set it apart from other controlled release delivery systems.

The support's chemical makeup

The drug's physical condition

The matrix's geometry,

The volume change over time

The release kinetic model

The drug is distributed in a swellable hydro-philic polymer, insoluble matrix of hydro-phobic materials, & plastic mate-

rials to regulate release of drug with varying solubility qualities.<sup>30</sup>

### Conclusion

In conclusion, oral extended and sustained release drug delivery systems play a crucial role in improving therapeutic outcomes by maintaining consistent drug levels in the body over an extended period. These systems enhance patient compliance by reducing dosing frequency and minimizing side effects associated with peak drug concentrations. Despite challenges in formulation and drug stability, ongoing advancements in pharmaceutical technology continue to improve the design and performance of these dosage forms.

**ACKNOWLEDGMENT:** The authors are highly thankful to the management of Sanskar Educational Group for constant support.

**Conflict of interest:** Nil



## Journal of Advanced Pharmaceutical Sciences and Natural Products

### REFERENCES

1. Ali J, Khar RK, Ahuja A. A Textbook of Biopharmaceutics & Pharmacokinetics. Birla Publications Pvt. Ltd. Delhi. 2008; 252-72.
2. Anwar Ma'ali; Naseef H, Qurt M, Abukhalil AD, Rabba AK, Sabri I. The Preparation and Evaluation of Cyanocobalamin Mucoadhesive Sublingual Tablets. *Pharmaceuticals*. 2023; 16(10): 1412.
3. Basak SC, Reddy JBM, Lucas Mani KP. Formulation and release behaviour of Extended Release Cyanocobalamin hydrochloride HPMC Matrix tablet. *Indian Journal of Pharmaceutical Sciences*. 2006; 68(5): 594-598.
4. Brahmankar HA, Jaiswal SB. Biopharmaceutics and Pharmacokinetics: A Treatise. VallabhPrakashan. 2000: 337, 348-357.
5. Borguist P, Korner A, Larsson A. A model for the drug release from polymeric matrix tablets: effect of swelling and dissolution. *J Controlled Release*. 2006; 113(3): 216-225.
6. Choudhari P. Formulation and Evaluation of Prolong Release Tablet of Antihypertensive Drug. *Research Journal of Pharmaceutical Dosage Forms and Technology*. 2018; 10(2): 55-59.
7. Chandran S, Laila FA, Mantha N. Design and evaluation of Ethyl Cellulose Based Matrix Tablets of Ibuprofen with pH Modulated Release Kinetics. *Indian Journal of Pharmaceutical Sciences*. 2008; 42(5): 234-240.
8. Chen X, Wen H, Park K. Challenges and new technologies of oral controlled release. *Oral Controlled Release Formulation Design and Drug Delivery: Theory to Practice*. 2010; 257-277.
9. Chien YW. Rate controlled drug delivery systems. 2nd ed. Marcel Dekker; New York, Revised and expanded. 2005; 2: 1-10.
10. Chien YW. Oral drug delivery systems in novel drug delivery pharmaceutical technology. Marcel Dekker Inc. New York, Basel. 1992; 152-96.
11. Divya B, Sreekanth J, Satyavati D. Formulation and evaluation of extended release matrix tablets of tenatoprazole sodium using synthetic polymers. *Journal of Young Pharmacists*. 2020; 12(2s): s39.
12. Eisai Submits New Drug Application for Mecobalamin Ultra-High Dose Preparation as Treatment for Amyotrophic Lateral Sclerosis in Japan. (PDF) Eisai.com. Retrieved 28 January 2018.
13. Gothi GD, Parinh BN, Patel TD, Prajapati ST, Patel DM, Patel CN. Study on Design and Development of Extended Release Tablets of Metoprolol Succinate. *Journal of Global Pharma Technology*. 2010; 2(2): 69-74.
14. Gunjal PT, Shinde MB, Gharge VS, Pimple SV, Gurjar MK, Shah MN. Design, Development and Optimization of S (-) Atenolol Floating Extended Release Matrix Tablets Using Surface Response Methodology. *Indian J Pharm Sci*. 2015; 77(5): 563-572.
15. KarishmaHalatwala\*, NayanRatnakar, Tushar Patel, Formulation, Optimization



## Journal of Advanced Pharmaceutical Sciences and Natural Products

- and Evaluation of Extended Release Tablet of Pregabalin for the Treatment of Diabetic Neuropathy *J Pharm SciBioscientific Res.* 2016 6(3):453-460.
16. Jaimin J Patel\*, Ravi R Patel, L.D. Patel, Yogesh Patel, Anil Raval. FORMULATION AND EVALUATION OF TIME DELAYED RELEASE TABLET OF CARVEDILOL PHOSPHATE. *Pharmacophore.* 2016; 7(2): 124-131.
17. James S, James C. Boylan *Encyclopaedia of Pharmaceutical Technology.* 4th ed. Marcel Dekker; 1997: 304-307.
18. Muhammad Asif Khan, Muhammad Saeed, Design, formulation, optimization and evaluation of sustained release tablets of domperidone *African Journal of Pharmacy and Pharmacology Vol. 5(16),* pp. 1882-1887.
19. KarishmaHalatwala\*, NayanRatnakar, Tushar Patel. Formulation, Optimization and Evaluation of Extended Release Tablet of Pregabalin for the Treatment of Diabetic Neuropathy. *J Pharm SciBioscientific Res.* 2016; 6(3): 453-460.
20. Nabin Karnal \*, Biswajit biswall , BhaveshBhavsar Formulation, Optimization and Evaluation of Extended Release tablets of Levetiracetam *Int.J.Pharm Tech Res.* 2014,6(2),pp 476-486.
21. Koyama K, Ito A, Yamamoto J, Nishio T, Kajikuri J, Dohi Y, et al. Randomized controlled trial of the effect of short term co-administration of Cyanocobalamin and folate on serum ADMA concentration in patients receiving long term haemodialysis. *American Journal of Kidney Diseases.* 2010; 55: 1069-1078.
22. Lachman L, Liberman AH, Kanig LJ. *The Theory and Practice of Industrial Pharmacy.* 3rd ed. Lea &Febiger; 1986: 430-456, 317-324.
23. Ma'ali A, Naseef H, Qurt M, Abukhalil AD, Rabba AK, Sabri I. The Preparation and Evaluation of Cyanocobalamin Mucoadhesive Sublingual Tablets. *Pharmaceuticals.* 2023; 16(10): 1412.
24. Mhaske NS, Kumar SS. Applications Of Sustained Release Dosage Form For Neuro Disorder: An Overview. *Journal of Pharmaceutical Negative Results.* 2022; pp. 6436-6446.
25. McDowell LR. *Vitamin in Animal and Human Nutrition.* Booksgoogle.com. ISBN 9780813826301. Retrieved 28 January 2018.
26. Muhammad Asif Khan, Muhammad Saeed. Design, formulation, optimization and evaluation of sustained release tablets of domperidone. *African Journal of Pharmacy and Pharmacology.* 2011; 5(16): 1882-1887.
27. Nava-Ocampo AA, Pastrak A, Cruz T, Koren G. Pharmacokinetics of high doses of cyanocobalamin administered by intravenous injection for 26 weeks in rats. *ClinExpPharmacol Physiol.* 2005; 32: 13-18.
28. Nishihata T, Tahara K, Yamamoto K. Overall mechanisms behind matrix Extended Release (SR) tablets prepared with hydroxypropyl cellulose 2910. *J Controlled Release.* 1995; 35: 59-66.
29. Pandey H, Tiwari VK, Prajapati SK. Extended Release bilayered tablets of Domperidone Maleate using hydrophilic



## **Journal of Advanced Pharmaceutical Sciences and Natural Products**

matrix system. *Indian Drugs*. 2007; 44(4): 261-263.

30. Patel, N., Akhtar, S., Vaibhav, B., Tiwari, S. Formulation, optimization and evaluation of extended release tablets of levetiracetam. *Int. J. Pharm Tech Res*. 2014; 6(2): 476-4