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Review Article

Transdermal Nanoparticle Patches: A New Era in Controlled Drug Delivery

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ABSTRACT

To get around the challenges of oral medicine delivery, the transdermal drug delivery method was developed. A transdermal patch is an adhesive pharmaceutical patch applied to the skin that allows a prescribed dosage to enter the bloodstream through the skin. This frequently aids in the recovery of a damaged body part. The preparation techniques for several transdermal patch types, including matrix patches, reservoir type, membrane matrix, drug-in-adhesive patches, and micro reservoir patches, are covered in this review article. Additionally, a study of the several transdermal dosage form evaluation techniques has been conducted. Transdermal drug delivery is a very appealing and cutting-edge method of getting medications through the skin since it can have a systemic effect. Given the current situation, producing safe medications with fewer harmful side effects associated with the majority of pharmacologically active substances requires particular attention. Transdermal drug delivery is a key paradigm that offers patients convenience, avoidance of first-pass hepatic metabolism, local targeting, and a decrease in toxic effects associated with a variety of drug classes, including analgesics, anti-inflammatory, antibiotics, antivirals, anaesthetics, and anticancer medications. The highly ordered structure of the skin, which serves as the primary barrier to drug penetration via the skin, makes even this route difficult. Nowadays, transdermal patches are widely employed as topical, transdermal, and cosmetic delivery systems. These patches are a major result of the advancements in skin science, technology, and expertise that have been made possible by clinical observation, trial-and-error, and evidence-based research that goes all the way back to the first human records. This review starts with the first topical treatments and tracks topical distribution to the contemporary transdermal patches, outlining the early trials, tools, and drug delivery systems that support the active ingredients in transdermal patches today. This is followed by consideration of the evolution in the various patch designs and their limitations as well as requirements for actives to be used for transdermal delivery. The properties of and issues associated with the use of currently marketed products, such as

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variability, safety and regulatory aspects, are then described. Their view concludes by examining future prospects for transdermal patches and drug delivery systems, such as the combination of active delivery systems with patches, minimally invasive microneedle patches and cutaneous solutions, including metered-dose systems.

INTRODUCTION

To provide a precise dosage of medication through the skin and into the bloodstream, a transdermal patch is utilised. The FDA initially authorised transdermal patches in 1981. There are currently transdermal delivery systems that contain nicotine to help people quit smoking, fentanyl for chronic pain, clonidine and nitroglycerin for cardiovascular illness, and scopolamine (hyoscine) for motion sickness. Transdermal delivery removes pulsed entry into the systemic circulation, permits continuous input of medications with brief biological half-lives, and offers controlled, continuous drug administration¹. Compared to traditional injection and oral techniques, TDDS has numerous benefits.. It lessens the strain that the oral route often puts on the liver and digestive system. It improves patient adherence and reduces negative pharmacological adverse effects brought on by transient overdose. It is particularly useful for patches that only need to be applied once a week. Patient adherence to medication therapy is facilitated by such a straightforward dose schedule².

The main components to a transdermal patch are:

- **Polymer matrix**—backbone of TDDS, which regulates the drug's release. The polymer should be nontoxic, chemically non-reactive, and not break down while being stored. It should also be reasonably priced. For instance, zein, gelatin, shellac, cellulose derivatives, gums, waxes, Silicon rubber, nitrile, acrylonitrile, polybutadiene, hydriin rubber, polyisobutylene, neoprene,

Polyacrylate, polyamide, polyurea, polyvinyl alcohol, polyvinyl chloride, polyethylene, polypropylene, polyvinylpyrrolidone, and polymethylmethacrylate³.

- **Drug-** For medications with the right pharmacology and physical chemistry, the transdermal route is a very alluring choice. Transdermal patches provide numerous medications that endure a lengthy first pass metabolism, medications with a limited therapeutic window, or medications with a brief half-life. such as nitroglycerine, fenatyl, etc.
- **Permeation enhancers-** Enhance the stratum corneum's permeability to get greater medication therapeutic concentrations. These come in three varieties: surface active, lipophilic solvent, and agents as well as two separate systems. For instance, DMSO⁴
- **Adhesive-** Increasing the stratum corneum's permeability will let the medication reach higher therapeutic levels.
- **Backing laminates-** should be highly flexible or have a low modulus. such as polyethylene and vinyl.
- **Release liner-** keeps the patch safe while being stored. Before using, the liner is taken out.
- Other excipients like plasticizers and solvents⁵.

Drug Delivery Routes across Human Skin

Drug molecules can penetrate by three pathways:

1. Sweat ducts
2. Hair follicles
3. Sebaceous glands

or

Directly across the stratum corneum.



Large, flat, polyhedral, plate-like envelopes filled with keratin—a substance derived from dead cells that have moved up from the stratum granulosum—make up the stratum corneum, the outermost layer of the epidermis. Dead cells devoid of nuclei make up the majority of this skin layer. New cells from the stratum germinativum (basale) continuously replace these dead cells when they slough off on the surface in the thin, air-filled stratum disjunctum. The thickness of the stratum corneum, which is made up of 10–15 layers of corneocytes, ranges from around 10–15 μm when dry to 40 μm when hydrated. It primarily consists of a multi-layered "brick and mortar" structure. In an intercellular matrix (mortar) made up of long chain ceramides, free fatty acids, triglycerides, cholesterol, cholesterol sulphate, and sterol/wax esters, the structure of keratin-rich corneocytes (bricks) is similar. Keratinocytes in the middle to upper stratum granulosum release their lamellar contents into the intercellular space, generating the intercellular lipid matrix. The

stratum corneum's first layers reorganise to create wide, intercellular lipid lamellae, which subsequently unite to form lipid bilayers. The lipid phase behaviour differs from that of other biological membranes due to the lipid content of the stratum corneum. A vital component of the stratum corneum, water serves as a plasticiser to keep it from cracking and contributes to the production of natural moisturising component that keeps the skin soft. Determining the main route of drug penetration within the stratum corneum is crucial to comprehending the physicochemical characteristics of the spreading drug and vehicle effect throughout the stratum corneum. A molecule moving across For the majority of medications, it is detrimental to diffuse into and across several hydrophilic and hydrophobic domains. As a result, it is presently thought that the intercellular route is the main mechanism that most medications pass through the stratum corneum⁶.

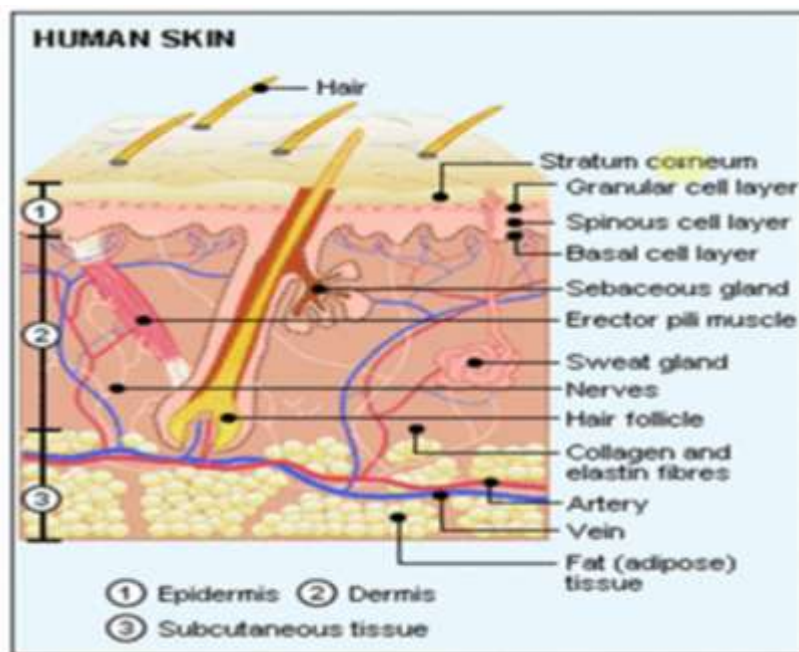


Fig. 1: Transverse section of skin showing routes of penetration 1. Through the sweat ducts; 2. Directly across the stratum corneum; 3. Via the hair follicles²

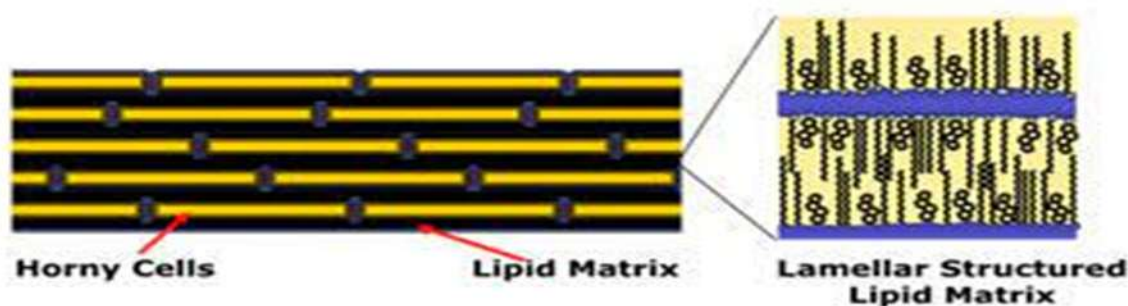


Fig. 2: Schematic structure of the stratum corneum according to the brick-and-mortar model³.

For more than ten years, transdermal medication delivery patches have been available on the market⁴. Over-the-counter nicotine patches that aid in quitting smoking are the most widely used transdermal drug delivery patches. The term "transdermal drug delivery patch" raises a number of queries, like what kind of material it is composed of, how it operates, and so on. Transdermal patches were initially developed to treat motion sickness in astronauts travelling into space⁷. However, a few drawbacks have prevented this medication delivery technology from reaching its full potential. The quantity of medications may be restricted by localised skin irritation and sensitisation. Effective transdermal medications possess molecular masses that are only a few hundred Daltons, which thus restricts the drug's dosage. Other drawbacks include delayed absorption, high prescription costs, and challenges in administering hydrophilic medications⁸. For both local effects and systemic therapy, transdermal drug delivery systems are sustained and controlled release devices that deliver medications or other therapeutic agents, such as peptides, nucleic acids, etc. via application on the skin's surface⁹. Although many transdermal systems with various medications have been developed, their practical application is restricted by the skin's penetration barriers. Many techniques have been developed and patented to aid with transdermal absorption¹⁰. These innovative technologies include, for example,

electroporation, iontophoresis, microneedles, ultrasound, and above all the application of nanotechnology¹¹. With an area of 1.5 to 2.0 m² in adults, the skin is the biggest organ in the human body by mass. Since the beginning of human history, drugs have been administered topically to cure surface conditions, transdermally to administer therapies to treat systemic illnesses, and as cosmetics. For example, ancient Egyptian and Babylonian medicine (c. 3000 BC) already widely used salves, ointments, potions, and even patches made of plant, animal, or mineral extracts. (Geller, 2010; Magner, 2005). However, it wasn't until the latter part of the 20th century that transdermal delivery methods became widely used, thanks to advancements in delivery technology that made it possible to administer medications precisely and consistently through the skin for systemic effects. The goal of every pharmaceutical firm and researcher is to create a safe and effective medication delivery system¹². Drug administration via the transdermal method can provide both systemic and local therapeutic effects¹³. Since transdermal drug delivery avoids gastrointestinal side effects and first pass metabolism, it is a desirable alternative to oral drug administration. Additionally, it can overcome the low patient compliance that is linked to other drug delivery methods¹⁴. Self-administered transdermal drug delivery enables the medication to enter intact skin over a predetermined amount of time to produce a local or systemic effect¹⁵.

Table 1. Drug product and clinical use of transdermal patches on the current market

Drug	Product name	Clinical use
Scopolamine	Transderm-Scop	Motion sickness
Nitroglycerin	Transderm-Nitro	Angina pectoris
Clonidine	Catapres-TTS	High blood pressure
Estradiol	Estraderm	Menopause
Fentanyl	Duragesic	Chronic pain
Nicotine	Nicoderm	Smoking cessation
Testosterone	Testoderm	Testosterone low level
Lidocaine/epinephrine	Iontocaine	Pain relief
Estradiol/norethidrone	Combipatch	Menopause
Lidocaine	Lidoderm	Pain relief
Norelgestromin	Ortho Evra	Contraception
Estradiol/levonorgestrel	Climara Pro	Menopause
Oxybutynin	Oxytrol	Overactive bladder
Lidocaine (ultrasound)	Sonol'trep	Pain relief
Lidocaine/tetracaine	Synera	Pain relief
Fentanyl HCl	Ionsys	Postoperative pain
Methylphenidate	Daytrana	ADHD
Selegiline	Emsam	Depression
Rotigotine	Neupro	Parkinson's disease
Rivastigmine	Exelon	Dementia

1. TDDS Classification Based on Their Technical Sophistication:

- Rate pre-programmed drug delivery system
- Activation modulated drug delivery system
- Feedback regulated drug delivery system
- Carrier based drug delivery system

a) Rate Pre-Programmed Drug Delivery System: Involves the creation of a system that administers medication by regulating the molecular diffusion of drug molecules within or around the delivery system across the eIt pidermal barrier¹⁶.

i. Polymer membrane permeation-controlled drug delivery system: Includes the system where the medication is contained inside a medication reservoir .This is taken care of by the polymer's semipermeable barrier, which controls release and has a particular Permeability¹⁷. There is some possibility development using the membrane process Microporous membranes allow for penetration. Stomach fluid resistance intestine targeted controlled release gastrointestinal device,

gel diffusion-controlled medication delivery system, and permeation controlled gastrointestinal delivery device.

ii. Polymer matrix diffusion-controlled drug delivery system : It is created by uniformly spreading drug particles within a rate-controlling carrier matrix. For instance, NitroDur. It provides a consistent transdermal infusion of nitroglycerine and is intended to be applied to intact skin for 24 hours¹⁸.

Microreservoir partitioned controlled drug delivery system : It uses high energy dispersion to distribute drug solution microparticles (which are aqueous in nature) throughout a polymer. For instance. The Syncromate implant was designed to administer Norgestomet¹⁹ subcutaneously.

b) Activation Modulated Drug Delivery System :This type of delivery system can be achieved by

i. Physical means:

Hydrodynamic pressure-controlled medication delivery system; osmotic pressure-activated drug delivery system.

- A drug delivery device that is activated by vapour pressure.
- A drug delivery mechanism that is mechanically actuated.
- A drug delivery method that is magnetically actuated.
- A drug delivery system that is electrically actuated.
- A drug delivery device that is activated by ultrasound.
- A drug delivery mechanism that is activated by hydration.
- **Chemical means:**
 - pH activated drug delivery system.
 - Ion activated drug delivery system.
 - Hydrolysis activated drug delivery system.
- **Biochemical means:**
 - Enzymes activated drug delivery system.
- **Feedback Regulated Drug Delivery System:** An agent that initiates the release of the drug, such as biochemicals, facilitates the release of the drug molecules from the transdermal system. In the body and is also controlled by its concentration via a feedback mechanism²⁰.
- Bio-erosion regulated drug delivery system.
- Bio-responsive drug delivery system.
- Self regulated drug delivery system¹⁷.

Carrier Based Drug Delivery System (Colloidal particulates carrier system): Hydrogels and other vesicular systems are involved microerythrocytes, microspheres, Aquasomes, dendrimers, transferosomes, etc²¹. They are favoured over the oral method of medication delivery because they systemic circulation due to a number of favourable factors;

2. ADVANTAGES OF TRANSDERMAL PATCHES:

- There is an improvement and an increase in bioavailability.
- Some patients have trouble swallowing pills and capsules.
- In order to facilitate swallowing, patients are tempted to smash medications, which eliminates the pills' controlled release properties.
- They are favoured over more uncomfortable hypodermic injections.
- Provide a risk of disease transmission and produce medical waste²².
- Increased patient cooperation because the procedure is easy, non-invasive, and practical, and there is more freedom in how medications are discontinued. through patch removal.
- Drugs delivered through the skin under control can offer less fluctuation and lower the concentration of the medication increase seen following the medications used orally²³.

3. DISADVANTAGES OF TRANSDERMAL PATCHES:

- The potential for local discomfort at the application location.
- The medication, the adhesive, or additional excipients in the patch formulation may result in erythema, irritation, and local oedema.
- Could result in allergic responses²⁴.
- It is necessary to have a molecular weight below 500 Da.
- A log P (octanol/water) of 1 to 3 is necessary for the permeate to cross the SC and underlying aqueous layers, indicating adequate aqueous and lipid solubility.

4. TYPES OF TRANSDERMAL PATCHES :



- **Single layer drug in adhesive patches:** A single layer of an adhesive polymer serves as a drug dispersion reservoir in Figure 1. The single layer is covered with an impermeable backing laminate. The medication is released from the backing laminate layer supporting the drug reservoir after being deposited in and adhering to the single polymer layer²⁵. One example of a single layer drug-in-adhesive transdermal patch that contains methylphenidate is the transdermal medication Daytrana®.
- **Multilayer drug in adhesive patches:** A drug reservoir layer and an adhesive layer with regulated medication release over time make up multilayer transdermal patches²⁶. Multilayer systems comprise both a permanent backing laminate and a temporary protective layer. Multilayer patches are used to administer hormone therapy, painkillers, and medications that promote quitting smoking; the duration of drug delivery can be up to seven days.
- **Vapour transdermal patches:** A single layer of sticky polymer with a vapour release feature that allows vapour to be released makes up vapour transdermal patches²⁷. There are several vapour dermal patches on the market, each with a distinct function. Nicoderm CQ®, for instance, are transdermal patches that contain nicotine vapour and essential oils that, when released, can aid in quitting smoking. In 2007, this product made its debut on the European market. Another kind of vapour patch that contains essential oils and can be used for decongestion is Altacura®. There are different kinds of vapour patches on the market that are used as sedatives or antidepressants²⁸ (Table I).
- **Membrane moderated transdermal reservoir patches:** A transdermal patch with a drug reservoir, an impermeable metallic plastic laminate backing layer, and a porous polymeric membrane that regulates drug release over time is seen in Fig. 1. Polymeric materials, such as ethylene vinyl acetate copolymer and hypoallergenic adhesive polymer, are used to make the membrane. The drug's molecular dispersion in a polymer matrix portion of the preparation regulates the drug's presence in the transdermal patch²⁹. Commercial transdermal patches with modified drug release include Catapres®, which contains clonidine for seven days, Transderm-Nitro®, which contains nitroglycerin for one day, and Transderm-Scop®, which contains scopolamine for three days (Table I).

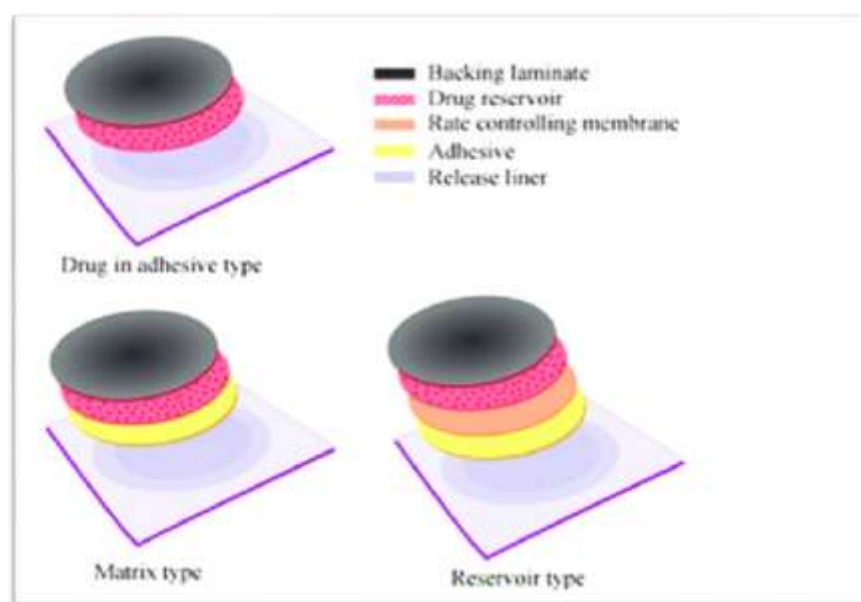


Fig. 1. Schematic diagram of various types of transdermal patches.

- **Microreservoir Transdermal Patches:**

Matrix dispersion and a drug reservoir are combined in microreservoir transdermal patches. The reservoir is made by spreading the drug suspension uniformly on a lipophilic polymer after it has been suspended in an aqueous solution of hydrophilic polymer³⁰. Thousands of tiny, insoluble spheres are formed

as a result of the strong shear mechanical stress used during dispersion. The drug level in the plasma is kept constant by the drug release profile, which adheres to a zero order rate of kinetic drug release. Since the medication dispersion must be thermodynamically stable, crosslinking polymeric agents are typically included³¹.

Table II. FDA approved other transdermal delivery systems

Drug	Product name	Transdermal delivery system
Flurandrenolide	Cordran® Tape	Transdermal tape
Testosterone	AndroGel®	Transdermal gel
Estradiol	Evamist®	Transdermal spray
Fentanyl HCl	IONSYS®	Iontophoretic patch
Insulin	Vyteris insulin patch®	Iontophoretic patch
Hydrocortisone	Tegaderm patch	Electrophotophoresis

- **Matrix system: drug-in-adhesive:** As illustrated in Fig. 1, the drug reservoir is made to disperse the medication on an adhesive polymer using single layer or multilayer transdermal patches. Either solvent casting or melting the sticky polymeric elements are used to deposit this drug-polymer matrix onto an impermeable backing layer³². There are several

commercial products of this kind of transdermal patch on the market. For instance, the NicoDerm® CQ transdermal patch contains nicotine to promote quitting smoking for up to 10 weeks, and the Climara® transdermal patch contains 100 microgrammes of oestradiol for one-day application (Table I).

- **Matrix systems: Matrix-dispersion:** A hydrophilic or lipophilic polymer matrix serves as the reservoir in a matrix transdermal patch, and the medicine is uniformly distributed throughout the matrix³³ by covering the plate with an impermeable laminate backing. A continuous medication flow through undamaged skin is provided by commercial matrix dispersion patch products like Nitro-Dur®, which comprises nitroglycerin and minitran (Table I).
- **Miscellaneous transdermal patches:** Transdermal patches with adhesive tapes, transdermal gel, transdermal spray, iontophoretic delivery, and photophoresis delivery are other FDA-approved transdermal matrix delivery methods, as indicated in Table II³⁴.

5. Evaluation Parameters:

1. Thickness of the patch: Using a digital micrometre, the thickness of the drug-loaded patch is measured at several sites. To guarantee the thickness of the prepared patch, the average thickness and standard deviation are calculated. Transdermal film thickness is measured at several locations on the film using a micrometre, screw gauge, or travelling microscope dial gauge³⁵.

2. Weight uniformity: Before testing, the created patches are dried for four hours at 60°C. A predetermined patch area needs to be chopped into several pieces and weighed using a digital balance. The individual weights must be used to get the average weight and standard deviation values³⁶.

3. Folding endurance: It is necessary to cut an even strip of material and fold it repeatedly at the same spot until it breaks. The folding endurance of a film is determined by how many times it can be folded in the same spot without breaking³⁷.

4. Percentage Moisture content: Each produced film must be weighed separately and stored for 24 hours at room temperature in a desiccator filled with fused calcium chloride. The films must be reweighed after 24 hours in order to calculate the percentage moisture content using the formula below³⁸.

$$\text{Moisture content (\%)} = \frac{\text{Initial weight} - \text{Final weight}}{\text{Initial weight}} \times 100\%$$

5. Content uniformity test: Ten patches are chosen, and each patch's content is established. Transdermal patches pass the content uniformity test if the content of nine of ten patches is between 85% and 115% of the prescribed value, and one patch has at least 75% to 125% of the stated value. However, 20 more patches are examined for drug content if three of them contain content between 75% and 125%. The transdermal patches pass the test if the range of these 20 patches is between 85% and 115%³⁹.

6. Moisture Uptake: Films that have been weighed are stored for 24 hours at room temperature in desiccators. After that, they are removed and placed in desiccators with a saturated potassium chloride solution at 84% relative humidity until their weight remains constant. The percentage of moisture uptake is computed as follows:⁴⁰

$$\text{Moisture content (\%)} = \frac{\text{Initial weight} - \text{Final weight}}{\text{Initial weight}} \times 100\%$$

7. Drug content: A certain volume of a patch must be dissolved in an appropriate solvent. After that, the solution must be filtered through a filter medium, and the drug content must be examined using the appropriate technology (either the UV or HPLC method). The average of three distinct samples is shown by each value⁴¹.

8. Tensile strength: A tensile tester (Shimadzu Autograph AG-X, Japan) assessed the patches'



tensile strength. The tested patches were positioned between cell grips after being trimmed to a size of $4 \times 1 \text{ cm}^2$. The film was subjected to force until it broke⁴². The dial reading in kilogrammes was used to calculate the patches' tensile strength. The following formula was used to determine the patches' tensile strength:⁴³

$$\text{Tensile strength} = \frac{\text{Tensile load at break}}{\text{Cross-section area}}$$

9. Shear Adhesion test: The purpose of this test is to determine an adhesive polymer's cohesive strength⁴⁴. The molecular weight, degree of cross linking, polymer composition, kind, and quantity of tackifier used can all have an impact. A stainless steel plate is covered with adhesive-coated tape, and to make the tape pull parallel to the plate, a certain weight is suspended from it. The time it takes to remove the tape from the plate is used to calculate the shear adhesion strength. The shear strength⁴⁵ increases as removal time increases.

10. Peel Adhesion test: Peel adhesion is the term used in this test to describe the force needed to remove an adhesive covering from a test substrate. The variables that influenced the peel adhesion qualities were the adhesive polymer's molecular weight and the kind and quantity of additives. The force needed to remove a single piece of tape is measured after it has been attached to a stainless steel plate or a preferred backing membrane. The tape is then lifted from the substrate at a 180° angle⁴⁶.

11. Water vapor transmission studies (WVT): Weigh one gramme of calcium chloride and put it

in previously dried, empty vials with the same diameter to determine WVT. Using an adhesive such as silicon adhesive grease, the polymer films are applied to the brim and left to set for five minutes. The vials are then precisely weighed and put in a humidity chamber that is kept at 68% relative humidity. The vials are measured once more at the conclusion of the first, second, and third days for a total of seven days in a row. A weight rise was regarded as a quantitative indicator of the amount of moisture that was transferred through the patch. In a different procedure that was published, vials containing 200 mL of saturated sodium bromide and saturated potassium chloride solution were placed in desiccators⁴⁷. The desiccators were sealed tightly, and a hygrometer was used to measure the humidity within. The process was then repeated after the weighed vials were put in desiccators. WVT is equal to W/ST . W is the weight increase during a 24-hour period; S is the exposed film area (cm^2); and T is the exposure time⁴⁸.

12. Rolling ball tack test: This test evaluates a polymer's tack-related softness. In this test, a 7/16-inch-diameter stainless steel ball is dropped upon an incline so that it rolls downward and encounters horizontal, upward-facing adhesive. Tack, which is measured in inches¹⁷, is determined by the distance the ball goes along the adhesive⁴⁹.

13. Quick Stick (peel-tack) test: In this test, the tape is dragged 12 inches per minute away from the substrate at 90° degrees Celsius. Tack value, which is measured and documented as the peel force necessary to break the adhesive-substrate bond, is given in ounces or grammes per inch width⁵⁰.

Table 3: : Marketed Products of Transdermal Drug Delivery System

S. No	Product	Active drug	Type of transdermal patch	Purpose
1.	Estraderm	Estradiol	Membrane	Postmenstrual syndrome
2.	Duragesic	Fentanyl	Reservoir	Pain relief patch
3.	Transderm-Scop	(Scopolamine)		Motion sickness
4.	Alora	Estradiol	Matrix	Postmenstrual Syndrome
5.	Climara	Estradiol	Matrix	Postmenstrual Syndrome
6.	Androderm	Testosterone	Membrane	Hypogonadism in males
7.	Captopress TTS	Clonidine	Membrane	Hypertension
8.	Combipatch	Estradiol	Matrix	Postmenstrual Syndrome
9.	Esclim	Estradiol	Matrix	Hormone replacement therapy
10.	Deponit	Nitroglycerine	Drug in adhesive	Angina Pectoris
11.	FemPatch	Estradiol	Matrix	Postmenstrual syndrome
12.	Lidoderm	Lidocaine	Drug in adhesive	Anesthetic
13.	Ortho Evra	Estradiol	Drug in adhesive	Postmenstrual Syndrome
14.	Testoderm TTS	Testosterone	Reservoir	Hypogonadism in males
15.	Habitraol	Nicotine	Drug in adhesive	Smoking Cessation
16.	Prostep	Nicotine	Reservoir	Smoking Cessation
17.	Nicotrol	Nicotine	Drug in adhesive	Smoking Cessation
18.	Vivelle	Estradiol	Reservoir	Postmenstrual syndrome
19.	Matrifen [®]	Fentanyl	Reservoir	Pain relief patch
20.	NuPatch 100	Diclofenac diethylamine	Drug in adhesive	Anti Inflammatory
21.	Nicoderm CQ	Nicotine	Drug in adhesive	Smoking Cessation
22.	Vivelle-Dot	Estradiol	Reservoir	Postmenstrual syndrome
23.	Minitran	Nitroglycerine	Drug in adhesive	Angina Pectoris
24.	Nitrodisc	Nitroglycerine	Micro reservoir	Angina Pectoris
25.	Nitroderm	Nitroglycerine	Matrix	Angina Pectoris
26.	TransdermNitro	Nitroglycerine	Reservoir	Angina Pectoris
27.	Oxytrol [®]	oxybutynin	Matrix	Overactive bladder
28.	Nuvelle TS	Estradiol	Drug in adhesive	Hormone replacement therapy
29.	Fematrix	Estrogen	Matrix	Postmenstrual syndrome
30.	Climaderm	Estradiol	Matrix	Postmenstrual syndrome

CONCLUSION:

Since 1981, transdermal drug administration systems have been employed as secure and efficient medication delivery tools. In the field of transdermal patches, significant advancements have been made. Many researchers are interested in the Transdermal Drug Delivery System because of its many benefits. Nowadays, a lot of new research is being done to use this system to incorporate newer medications. Transdermal dose forms might give doctors the chance to give their patients more treatment choices so that their care is optimised. Our knowledge of the nature of the stratum corneum barrier and how chemicals interact with and affect its structure has improved recently thanks to the application of several biophysical approaches. The establishment of structure-activity connections for enhancers and a deeper comprehension of how enhancers interact with the stratum corneum will help designers create enhancers with the best properties and the least amount of toxicity. Important aspects about transdermal drug delivery systems and their

evaluation procedure are covered in full in this article.

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