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م.م. عمار عبدالمجيد أحمد	اسم التدريسي
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Pharmaceutical Dosage forms and Drug Delivery Systems By Haward A. Ansel; latest edition.	المصادر والمراجع
Sprovel's American Pharmacy.	

محتوى المحاضرة

Suspension Dosage form

- Solutions
- Dispersions

Colloidal Dispersion: 1 nm to 0.5 μm

Fine Dispersion: 0.5 to 10 μm

Coarse Dispersion: 10 to 50 μm

Coarse Dispersion (10 to 50 μm)

- Emulsion
 - Suspension
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Differences in term of Separation

A pharmaceutical suspension is a coarse dispersion in which insoluble particles, generally greater than 1 μm in diameter, are dispersed in a liquid medium, usually aqueous.

Suspensions: preparations containing finely divided drug particles (the suspensoid) distributed somewhat uniformly throughout a vehicle in which the drug exhibits minimum degree of solubility.

Suspensions

- Some suspensions are available in **ready-to-use form**, i.e., already distributed through a liquid vehicle with or without stabilizers and additives.
- Other preparations are available as **dry powders** intended for suspension in liquid vehicles.

✓ Drugs unstable for long periods in aqueous vehicles (e.g., many antibiotics) are supplied as dry powders for reconstitution at dispensing.

Uses of Aqueous Suspensions

- Most commonly used for **oral administration** of insoluble drugs.
 - May also be used for **parenteral, topical, ophthalmic** administration.
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Important Note

- Complete and uniform redistribution of the dispersed phase is essential for accurate dosing.
 - Achieved by moderate agitation of the container.
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Reasons for Suspensions – Advantages

- Stability

- Taste masking
- Ease of swallowing
- Solubility
- Bioavailability

Disadvantages

- Physical instability
 - Bulky
 - Aesthetic problems
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Features Desired in a Pharmaceutical Suspension

- Settle slowly and redisperse easily (stable).
 - Particle size remains fairly constant.
 - Suitable viscosity → pour readily and evenly.
 - Acceptable taste, odor, color.
 - Must not decompose or support microbial growth during storage.
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Classification of Suspensions

Based on Classes

- Oral
- External
- Parenteral

Based on Particle Size

- Colloidal
- Coarse
- Nanosuspension

Based on Proportion of Solids

- Diluted
- Concentrated

Based on Electrokinetic Nature

- Flocculated
- Deflocculated

Properties of Suspension

- Sedimentation behavior
- Electrokinetic properties

Factors affecting stability:

- Particle size
- Wetting of particles (surfactant/wetting agent)
- Particle movement / viscosity / sedimentation
- Electrical charge (zeta potential)
- Flocculation vs deflocculation
- Brownian motion

1. Sedimentation behavior

Settling of particles/floccules occurs under gravity.

Stokes' Law

Sedimentation rate \propto square of particle diameter.

- Ideal particle size: 1–50 μm .
- Avoid too large particles:
 - Rapid sedimentation, unstable

- 5 μm : gritty, cause irritation (eye/injection)
 - 25 μm : may block needles
 - Avoid too small particles:
 - Tend to form **compact cake** difficult to redisperse
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Particle Size Reduction

- Reduction improves stability (slows sedimentation).
 - Achieved by:
 - **Micropulverization** (10–50 μm , oral/topical).
 - **Jet milling / micronizing** (<10 μm , parenteral/ophthalmic).
 - **Spray drying** → very fine powders via drying drug solution with warm air.
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2. Density of particles & medium

- Particles denser than aqueous vehicle → faster settling.
 - Less dense particles → float, hard to redisperse.
 - If equal → suspension more stable.
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3. Viscosity of dispersion medium

- Higher viscosity reduces sedimentation.
- But too high viscosity → poor pouring & redispersion.
- Controlled by vehicle + solid content.
- Often improved with **suspending agents**:
 - Carboxymethylcellulose (CMC)
 - Methylcellulose

- Microcrystalline cellulose
 - Polyvinylpyrrolidone (PVP)
 - Xanthan gum
 - Bentonite
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Effect of Particle Shape on Stability

- **Needle-shaped particles** → form hard, non-redispersible cake.
 - **Barrel-shaped particles** → more stable.
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Solid Particle – Liquid Interactions

- Solubility requires chemical similarity.
 - Many modern hydrophobic drugs → poor water solubility → formulated as suspensions.
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Electrical Double Layer Theory

- Hydrophobic drugs in water acquire charge due to ionization of water.
 - OH^- adsorbs on particle surface → negative charge.
 - Two layers form:
 - **Fixed layer** (Stern) – tightly bound ions.
 - **Diffuse layer** (Gouy–Chapman) – mobile ions.
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Zeta potential (ζ): potential difference between medium & stationary fluid layer on particle.

Zeta potential & colloid stability:

- 0 to ± 5 mV → rapid coagulation.
- ± 10 to ± 30 mV → incipient instability.
- ± 30 to ± 40 mV → moderate stability.

- ± 40 to ± 60 mV \rightarrow good stability.
 - ± 61 mV \rightarrow excellent stability.
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Effect of Excipients on Double Layer

- Electrolytes (e.g., NaCl) increase mobile charges.
 - Surfactants change charge magnitude and sometimes its sign.
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Particle Interactions

1. Van der Waals attraction
 2. Electrostatic repulsion
- Higher repulsive force (zeta potential around ± 30 mV) \rightarrow particles remain apart \rightarrow stable suspension.

