

SEGREGATION



Dr.D.VARUN
Professor & Academic Director
SRI INDU INSTITUTE OF PHARMACY
Hyderabad

SEGREGATION

- DEMIXING / POWDER SEGREGATION
- REQUIREMENTS FOR SEGREGATION
- PARTICLE PROPERTIES THAT PRONE TO SEGREGATION
- APPROACHES TO RECTIFY SEGREGATION
- ORDERED MIXING
- SEGREGATION IN ORDERED MIXING

POWDER SEGREGATION (DEMIXING)

- Particulate solids tend to segregate by virtue of difference in size,density.shape...
- Occurs during mixing as well as handling of completed mix, more in free flow Powders
- More severe with free flowing, cohesionless / nearly cohesionless particles
- Segregation attributed to various types of mixers,
 - Principally convective motion , non-segregating
 - shear / diffusive mixing,segregating

- Opposite to mixing
- Imp in pharma pdt's –random to non random, or none
- Care to taken during handling (later mixing of powders) transfer to filling machines / hopper in tab, capsule..
- Cause in increase in content variation in samples of mix, fail
- If this occurs in filling machine., variation in weight
- Mainly due to monosized
- To some extent in powder bed ..vibration, or when great flowability

REQUIREMENTS FOR SEGREGATION

- Necessary and sufficient conditions.....
 - ✧ various mixture components exhibit mobilities for Interparticulate relative displacement, which differ.
 - ✧ mixture faces a field which exerts a directional motive force on particles. or a gradient capable of inducing / modifying interparticulate movement
- Requirements for segregation can arise in many ways....
 - ✧ differences in mixture component mobilities
different sizes, densities and surface characteristics
 - ✧ earth's gravitational force, centrifugal, electrical magnetic field generated during processing

PARTICLE PROPERTIES THAT PRONE TO SEGREGATION

- Chances of segregation by.....
 - particle size effects
 - particle density effects
 - particle shape effects

PARTICLE SIZE EFFECTS

Percolation segregation....

- occur in static powders.....greater when powder bed dilates / disturbed
- Domestically in cereal packets, coffee jars..smaller congregate to bottom
- Powder bed disturbed.....
(vibration, stirring or pouring)

TRAJECTORY SEGREGATION

during mixing larger particles will have high K.E.,
Separation of particles of different sizes

TRAJECTORY SEGREGATION + PERCOLATION SEGREGATION

occurrence of larger particles at the edge of heap
when poured from container.

ELUTRIATION SEGREGATION / DUSTING OUT

small particles as dust due to air currents, after
discharge get sedimented over coarser particles

PARTICLE DENSITY EFFECTS

- ☞ more dense ..move downwards,even of similar size
- ☞ trajectory segregation also occur with diff densities
- ☞ If denser particles are smaller, density has an effect on Percolation segregation
- ☞ Size and density effects cancel each other..when larger Particles are more dense.

PARTICLE SHAPE EFFECTS

- ☞ Spherical particles....more flow...easy mix..more segregate
- ☞ Irregular/needle shaped particles...interlocked,reduce the tendency to segregate
- ☞ Non spherical particles have surface area to wt ratio(specific surface area),decrease segregation by cohesive forces(greater contact area),also increase dusting out

APPROACHES TO RECTIFY SEGREGATION

- ✧ Selection of particle size fractions (sieving to remove fines)
- ✧ to achieve drugs and excipients of same narrow particle size range
- ✧ Milling of components, followed by sieving...at 30um , no Serious segregation
 - ✧ Controlled crystallization of drug/ Excipient for same range
- ✧ Selection of excipients similar to active components
- ✧ Granulation of powdered mix...evenly distributed reduce the extent of vibration
 - ✧ Limit the powder residence time
- ✧ Use equipment where no transfer of mass
(FBD , high speed mixer, granulator for mixing and Granulating)
- ✧ Production of an ordered mix

ORDERED MIXING

- Micronised particles ,adsorbed onto active sites of larger carrier particles....
 - this has an effect of minimising segregation while maintaining good flow properties.....ORDERED MIXING
- Mainly used in production of dry antibiotics...where water added
- Pharmaceutical mixes..partly ordered.partly random
- Degree of mixing superior to random mix
- Mainly in direct compressible formulations in prevention of
- Seggregation from directly compressible bases

SEGREGATION IN ORDERED MIXES

Carrier particle vary in sizes---

diff size particles..diff surface area to wt ratio,contain different Amounts of adsorbed material per unit area
drug areas where smaller carrier particles congregate...

.....**ORDERED UNIT SEGREGATION**

Competition for active sites on carrier molecules

displacement of adsorbed material,which may congregate
Owing to small size...

.....**DISPLACEMENT SEGREGATION**

Insufficient carrier molecules

excess material can't be absorbed and gets separated

..... **SATURATION SEGREGATION**