

Nanocarriers for targeted drug delivery

Benefits and challenges of nanotechnology for medicine

Russell Maguire

ENG14131 Advanced Semiconductor Devices

Durham University

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Nanocarrier delivery systems

Definition: nanocarrier

- Biocompatible nanoparticle encapsulating a drug

Characteristics

- Nanometre scale
- Hydrophilic
- Low toxicity
- Soluble

Targeted drug delivery

How is targeted drug delivery effective?

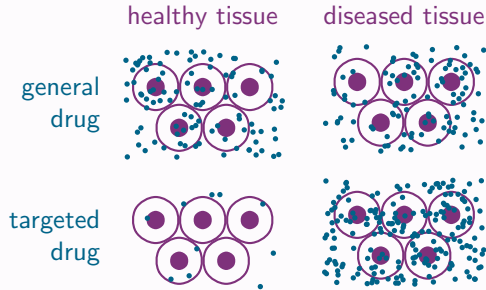


Figure 1: Goal is to increase drug concentration ratio for diseased tissue.

Targets diseased tissue

- Maximise therapeutic effects
- Minimise effective drug dose

Avoids healthy tissue

- Minimise side-effects

Vascular permeability

Definition: vascular permeability

- Tendency for molecules to leak from blood vessels into interstitial space between cells.

Contributing factors

- Permeability enhancers such as *bradykinin* and *nitric oxide*
- Dense and defective vascular network
- Enhanced vascular permeability has been observed in solid tumours and inflamed tissue [1]

[1] H. Maeda *et al.*, *Journal of controlled release* **65**, 271–284 (2000).

Enhanced Permeability and Retention effect

Definition: EPR effect

- Tendency for large molecules, such as nanocarriers, to accumulate in solid tumours and inflamed tissue

Accumulation procedure

- ① Molecules leak into interstitial space due to enhanced vascular permeability
- ② Smaller molecules diffuse back into blood vessels and lymph nodes
- ③ Larger molecules become trapped

Stimuli-responsive nanocarriers

Biological stimuli

- Tumours have been observed with lower pH than healthy tissue [2]
- Organelles maintain their own unique pH and redox status
- Difference in intracellular and extracellular redox status [3]

External stimuli

- Electromagnetic fields
- Heat
- Ultrasound
- Light

[2] L. E. Gerweck, K. Seetharaman, *Cancer research* **56**, 1194–1198 (1996).

[3] G. Saito *et al.*, *Advanced drug delivery reviews* **55**, 199–215 (2003).

Thank you, any questions?

References

1. H. Maeda *et al.*, *Journal of controlled release* **65**, 271–284 (2000).
2. L. E. Gerweck, K. Seetharaman, *Cancer research* **56**, 1194–1198 (1996).
3. G. Saito *et al.*, *Advanced drug delivery reviews* **55**, 199–215 (2003).