APPLICATION OF NANOTECHNOLOGY IN BRAIN TUMOR SPECIALLY GLIOBLASTOMA MULTIFORM

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GLIOBLASTOMA MULTIFORM...

Among all types of cancers, Glioblastoma Multiform (GBM) have one of the lowest survival rate. despite of treatment methods such as surgery, chemotherapy and radiotrary,

survival rate is between 12-15 month
5-y survival is 9.8%

 GBM: uncontrolled cell proliferation, high aggressive ability, high angiogenesis
, cellular genetic instability



MRI image of GBM

GBM...

□ major obstacles to treating GBM :

- 1. Feature of GBM
- 2. side effects and inefficacity of chemotherapeutic agents
- 3. blood-brain barrier (BBB)
- 4. blood-cerebrospinal fluid barrier (BCB)
- 5. drug resistance(...)
- 6. low bioavailability of drug
- 7. brain cancer stem cells (BCSCs)
- 8. Drug penetration into tissue is limited to $\sim 1 \text{ mm}$

BLOOD-BRAIN BARRIER...

The blood-brain barrier (BBB) is a major obstacle to treating GBM

The BBB is highly permeable to water, CO2, oxygen and lipid-soluble substances like alcohol. impermeable to plasma proteins

selective permeability of the BBB arises from the tight junction, efflux pumps, multidrug resistance proteins and degrading enzymes

The vast majority of GBMs recur within2 cm of the original tumor focus



STRATEGIES OF OVERCOMING THE BBB

□ Four major approaches:

1-Invasive approaches rely on the direct administration of drugs into the brain; catheter, drug implants, osmotic disruption of the BBB

2- Pharmacological approach relies on modification of drugs to enable penetration through the BBB

3- Physiological approaches; specific receptors or transport mechanisms in order to penetrate the BBB such as glucose, insulin, growth hormones and low-density lipoproteins, VEGF, low-density lipoprotein receptor proteins (LRPs)

Disadvantages: long recovery times, decreased drug uptake and low receptor mediated transport

NANOTECHNOLOGY AND NANOMEDICINE...

4- The final approach to bypass the BBB is through the use of NPs

Drug delivery: 76% share of scientific papers



□ In brain cancer research: with the aim of imaging and drug delivery solid-inorganic (magnetic Fe3O4 NPs, gadolinium NPs, gold NPs [AuNPs] and semiconductor quantum dots [QDs] and organic-based (dendrimer, hydrogel and polymer NPs

NPS AS 'THERANOSTICS' ...

□ It should be apparent that one of the most promising aspects of nanomedicine is the multi modality of a NP in any given application

Incorporation of both diagnostic and therapeutic modalities

Therapy Diagnosis Theranostic nanoparticle Imaging agent Therapeutic agen Therapeutic monitoring

Utilized polymer-coated magnetic NPs to both deliver the anticancer drug eprubicin and provide an MRI contrast agent in brain tissue

EFFECTS OF NP PROPERTIES FOR PENETRATION

1- size
2-shape
3-surface charge
4- surface coating
5- ...



www.pnas.org/cgi/doi/10.1073/pnas.1304504110

IN OUR STUDIES ...









SEM; MTX-PLGA

Powder; Curcumin-PLGA TEM; SLN- Indirubin

Albumn- MTX



SEM; Albumin - Imatinib



SEM; Albumin -Indirubin

