



Sperm cells as Targeted Delivery Agents in Tumor Therapy: A Comprehensive Review

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ABSTRACT:

Global cancer

related deaths continue being among the most prominent causes. Traditional drugs face serious disadvantages, particularly their lack of specificity, leading to unwanted side effects. Innovations in drug delivery recently gained more interest through 'biologically inspired' systems such as micromotors; one example can be a sperm cell and targeted drug delivery. The paper focuses on the application of sperm cells as natural micromotors for the targeted delivery of chemotherapeutic agents, mechanisms, efficacy, advantages, and future directions in clinical applications

Introduction :

Cancer treatment, especially regarding gynecological types such as ovarian and cervical cancer, has undergone dramatic changes for the past few decades. Still, the majority of patients undergo many side effects like other systemic therapies because of the non-specific distribution of drugs. It is this inducing structural deficit in the body for which the researchers couldn't find a very efficacious way of new systems for drug delivery directly targeting such an action for improved therapeutic outcome, lowering systemic toxicity. Sperm cells have introduced interesting possibilities as drug delivery vehicles because of their inherent motility and fitness to access highly complicated environments. They can thus be very usefully exploited to target tumors within the female reproductive tract, which has advantages in further refinement in specification as sperm cell biology enables them to efficiently navigate the female reproductive tract. The current review provides insights into some recent investigations with sperm cells toward targeted drug delivery and clinical possibilities.

2. Sperm-Based Drug Delivery Mechanism :

2.1 Loading Mechanism

Loading sperm cells with chemotherapeutic agents is recently proven to be feasible. The known anticancer drug, DOX-HCl, was successfully encapsulated into the human sperm head incubated with sperm cells in particular by loading mechanism. DOX-HCl can get absorbed passively into sperm head. Approximately 5.3 pg of DOX-HCl can be loaded without affecting sperm motility for each sperm cell.

2.1.1 Encapsulation Techniques

Various techniques have been developed to enhance loading efficiencies of the drugs in spermatozoa. Some of the techniques entail electroporation: a technique of introducing drugs into spermatozoa through pulses of short electric pulses which open the pores on the side of the sperm membrane long enough for the larger drug molecules to pass through. Nanocarriers may also be used to shield the drug during transport and to ensure its site-specific release.

2.2 Tumor Localization

During loading, sperm cells move through a body utilizing their natural motility to finally aggregate near tumor sites. Their chemotactic capacity-which means that the ability of sperm cells to respond to chemical gradients-can be enhanced to ensure greater tumor targeting. The sperm cells can be controlled with external stimuli such as magnetic fields, and it will ensure that the sperm cells reach the tumor area in the proper sequence.

2.2.1 Advanced Steering Mechanisms

Materials science advances led to sophisticated systems steering sperm cells toward target sites with more direct targeting. The steering systems can be nanoparticle-labeled inside the sperm or an external magnetic field to orient or drive sperm movement toward focusing or enhancing delivery efficiency toward target sites.

3. Tumor Targeting Application

3.1 Ovarian and Cervical Cancer

The research area for drug delivery systems using sperm cells has been greatly examined, particularly focusing on gynecological cancers. Ovarian cancer is considered to be a poor prognosis and even difficult-to-detect disease. Recent reports have demonstrated the cytotoxic effects of DOX-HCl-loaded sperm cells on 3D ovarian cancer cell cultures obtained from patients, resulting in vast reductions in cell viability.

3.1.1 Mechanistic Insight

The cytotoxicity of the sperm-mediated delivery could be through both direct action of the chemotherapeutic agent and the physical disruption caused by the motility of the sperm. The microenvironments created by the sperm cells enhance penetration of the drugs into the targeted elements and can increase contact with tumor cells. This partially may account for enhanced efficacy compared with conventional drug delivery technologies.

3.2 Other Types of Cancers

Much interest has been gaining in other cancers while more concentration is being given to gynecological cancers. In the male reproductive system, prostate and testicular cancers similarly originate from malignancies; therefore, sperm-mediated drug delivery may be utilized by utilizing sperm cells for these cancers as well. Tumors which anatomical places are very difficult to target are also very good candidates because of the ability of sperm to penetrate through complex biological fluids.

3.2.1

In addition to Cancer Therapies Besides Cancer Sperm Cells may be used in the treatment of other diseases as drug carriers. Considering that sperm cells can fuse with the majority of cell types, targeted delivery of gene therapies or vaccines can be applied, and therefore, a new treatment method for numerous diseases is opened.

4. Benefits and Drawbacks

4.1 Benefits

The advantage in applying sperm cells as drug carriers includes the following:

- **•Biocompatibility:** Since sperm cells are naturally found in the system, they are less likely to induce an immune response compared to synthetic carriers.
- **•Natural Motility:** The natural motility properties of the sperm have the ability to swim automatically and in a good direction through biological fluids, thereby reaching target sites more efficiently.
- **•Cell Fusion Capabilities:** Sperm cells have a unique property of being able to fuse with somatic cells, which may increase drug uptake in targeted cancer cells

4.2 Limitations

Despite promises, many hurdles are in the way:

- **•Control over Navigation:** The external guidance mechanisms look promising, but precise control of sperm movement in vivo is still an important challenge
- **•Off-Target Effects:** Interaction between sperm cells and non-cancerous cells has to be properly ruled out because it may cause some unwanted effects
- **•Regulatory Barriers:** Sperm-based therapies may pose particular regulatory problems with safety and effectiveness assessments for clinical applications
- **Ethical Considerations:** There are ethical issues regarding the consent and clinical potential misuse of human sperm.

5. Conclusion :

Future experiments should achieve best techniques when loading sperm for maximum drug encapsulation and potentially even more creative directions for the external guiding and control of the sperm. Additionally, multi-sperm carriers may allow effective delivery in that transport of over one sperm cells could result in a more general overall therapeutic effect in the body.

5.1 Preclinical studies

Suitable preclinical studies allow for the assessment of *in vivo* behavior of sperm-mediated drug delivery systems. In these, information critical to pharmacokinetics, biodistribution, and even effects on surrounding tissues can be obtained.

5.2 Combination Therapies:

Sperm-mediated delivery systems can be combined with other therapeutic modalities such as immunotherapy or radiotherapy at the site of tumor targeting to enhance the efficiency of treatment while reducing the side effects of therapies.

6. Conclusion :

Sperm cells represent a new and promising drug delivery strategy for the treatment of cancer therapy. Particularly, their biological features, so able to navigate complex scenarios, make them suitable carriers of chemotherapeutic agents, especially in gynecological cancers. Yet, there are still a number of researchers and technological developments that continue to intervene in changing paradigms in the treatment of cancer to help improve patient outcome.

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