□ Cancer Vaccine for Glioblastoma

A cancer vaccine for glioblastoma is an immunotherapy designed to stimulate the immune system to recognize and attack glioblastoma (GBM) tumor cells by targeting tumor-specific or tumor-associated antigens.

□ Rationale

Glioblastoma is a highly aggressive brain tumor with poor prognosis. Traditional therapies offer limited survival benefit. Cancer vaccines aim to:

- Induce T cell-mediated immune responses
- Promote immune memory against tumor recurrence
- Target specific tumor antigens or neoantigens
- Overcome glioma-associated immune suppression

☐ Types of Glioblastoma Vaccines

Туре	Description Example(s)		
Peptide Vaccines	Short tumor antigen fragments induce antigen-specific T cells	Rindopepimut (EGFRvIII)	
Dendritic Cell Vaccines	Patient DCs loaded with tumor lysate or peptides ex vivo	DCVax-L	
mRNA Vaccines	Encode neoantigens in mRNA to trigger T cell activation	NOA-16 Trial	
Neoantigen Vaccines	Personalized to each patient's unique tumor mutations	Moderna/BioNTech platforms	
Tumor Lysate Vaccines	Use whole tumor cell lysate to broaden antigen exposure	HSPPC-96 (heat-shock protein)	

☐ Target Antigens in GBM

- EGFRvIII mutant receptor in 25-30% of GBM
- IDH1 R132H mutation in lower-grade gliomas
- WT1, SOX2, Survivin tumor-associated antigens
- Personalized neoantigens identified through sequencing

☐ Key Clinical Trials

Trial Name	Туре	Target/Strategy	Phase	Outcome/Status
NOA-16	mRNA vaccine	Personalized IDH1 neoantigens	Phase I	Safe, immunogenic (93% T cell response)
AMPLIFY-NEOVAC		IDH1 vaccine + anti-PD- L1	Phase I	Ongoing

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Trial Name	Туре	Target/Strategy	Phase	Outcome/Status
DCVax-L	DC-based vaccine	Tumor lysate-loaded DCs	Phase III	Improved survival in long- term subgroup
Rindopepimut	Peptide vaccine	EGFRvIII	Phase III	No OS benefit → development discontinued

△ Challenges

- Antigen heterogeneity → not all tumor cells express the same target
- **Immune suppression** → GBM microenvironment inhibits T cells
- HLA restriction → some vaccines only work for patients with certain HLA types
- Time and cost of personalized vaccine design

□ Future Directions

- mRNA platforms for rapid personalization and multi-antigen delivery
- Al-based neoantigen prediction
- Combination therapies with:
 - 1. Checkpoint inhibitors (e.g., anti-PD-1)
 - 2. Oncolytic viruses
 - 3. Radiotherapy
- Local delivery methods (e.g., intratumoral injection, hydrogel implants)

□ Summary

Cancer vaccines for glioblastoma represent a promising class of immunotherapy with a growing body of early clinical evidence. Personalized mRNA vaccines and dendritic cell-based strategies are showing **immunogenicity and feasibility**, especially in combination with other treatments. Overcoming immune suppression and tailoring vaccines to tumor evolution are key to their future success.

□ Related Pages

- Glioblastoma
- Glioblastoma Immunotherapy
- NOA-16 Trial
- AMPLIFY-NEOVAC Trial
- Dendritic Cell Vaccines
- Personalized mRNA Neoantigen Vaccine
- Checkpoint Inhibitors

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Last update: 2025/03/26 06:07

