Therapy resistance



- Selection of resistance mutations
- Adaptive cellular reprogramming
- Lineage plasticity

Genetic selection



- Permissive mutational landscape
- Evolutionary bottlenecking
- Increasing aneuploidy

Plasticity



- Transcriptional heterogeneity
- Rewiring epigenetic landscape
- Metabolic adaptability

Macroenvironment systemic reprogramming



- Dysregulated systemic inflammation
- Pre-metastatic niche creation
- Cachexia

Principles of metastasis

Organ-specific microenvironment adaptation



- Reciprocal reprogramming of tumor and host
- Selection of organadapted transcriptional states



Dormancy

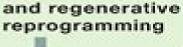


- Dynamic equilibrium with immune surveillance
- Reversible cancer cell quiescence





- Altered antigen presentation
- Dysregulated cytokine secretion
- Innate immunity suppression
- Impaired effector immune cell recruitment and cytotoxicity



Developmental



- Transient adoption of fetal-like multipotential states
- Reversion along developmental continuum

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