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The Role of Tuberin in DNA Damage Repair During Cell Proliferation

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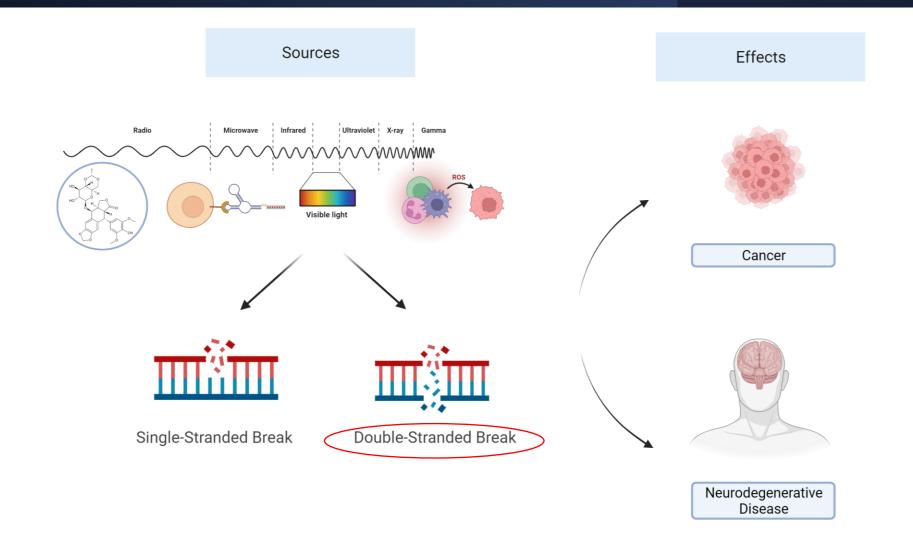
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The Role of Tuberin in DNA Damage Repair During Cell Proliferation

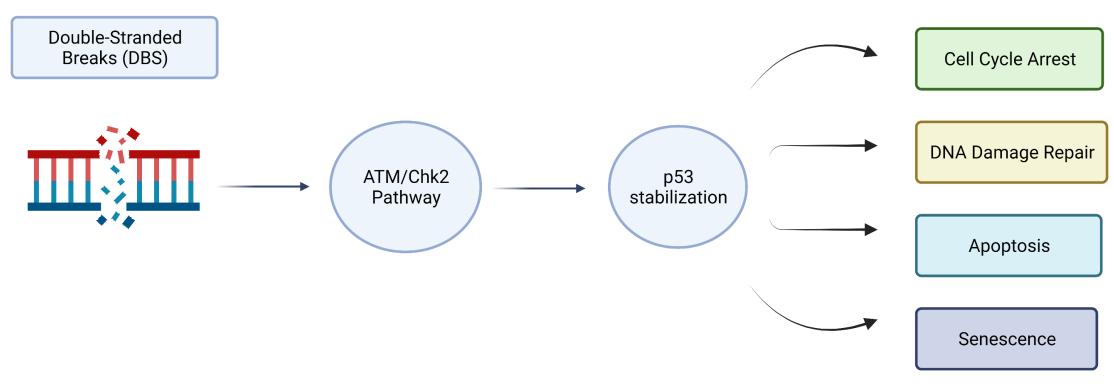
Kadila Adili

Why is DNA Damage Important?



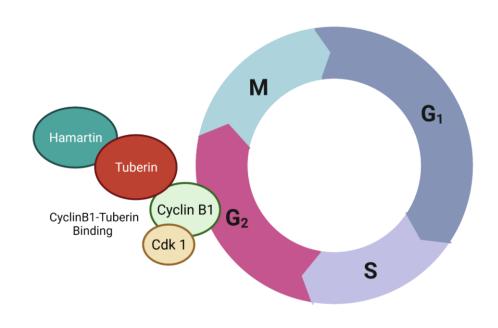


DNA Damage Response & p53





What Role Does Tuberin Play In The Cell Cycle?



Weak
CyclinB1-Tuberin
Binding

Cyclin B1

G2

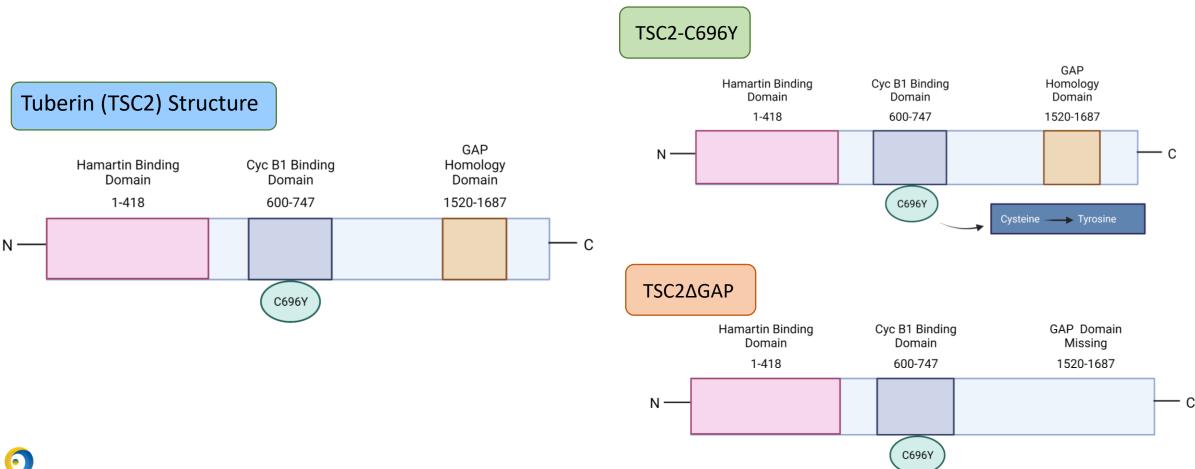
Hamartin
Tuberin

High Nutrient Conditions: Mitotic Delay & Increase in Cell Size

Low Nutrient Conditions: Progression into Mitosis

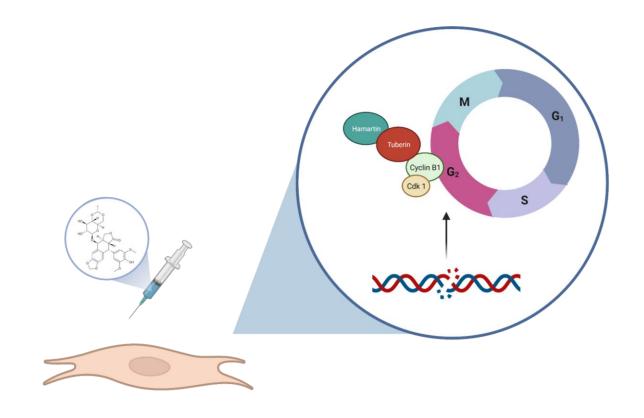


Tuberin Structure and Mutants



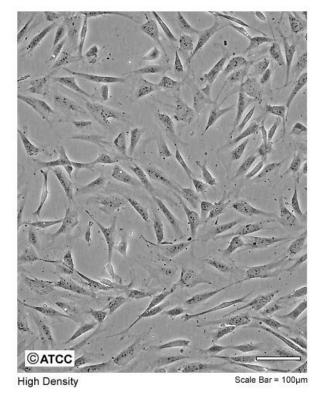
How does the Tuberin-Cyclin B1 complex play a role in DNA Repair?

Hypothesis: The Tuberin-Cyclin B1 complex plays a role in DNA damage repair through G2/M arrest.

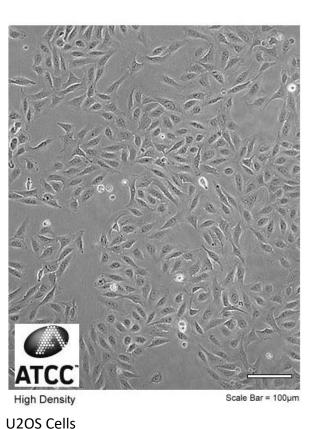


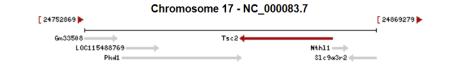


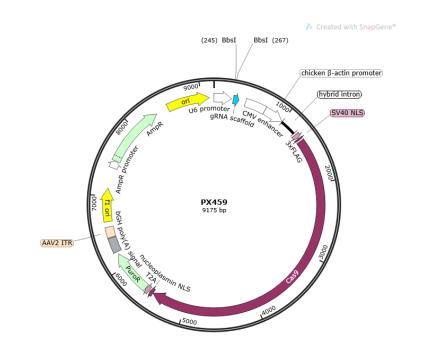
NIH/3T3 & U2OS Cell Line Rationale







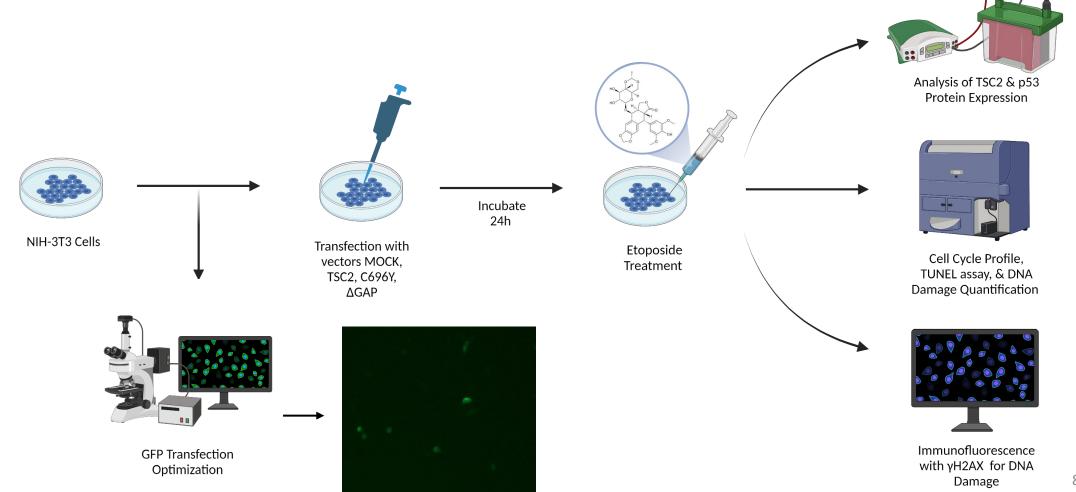






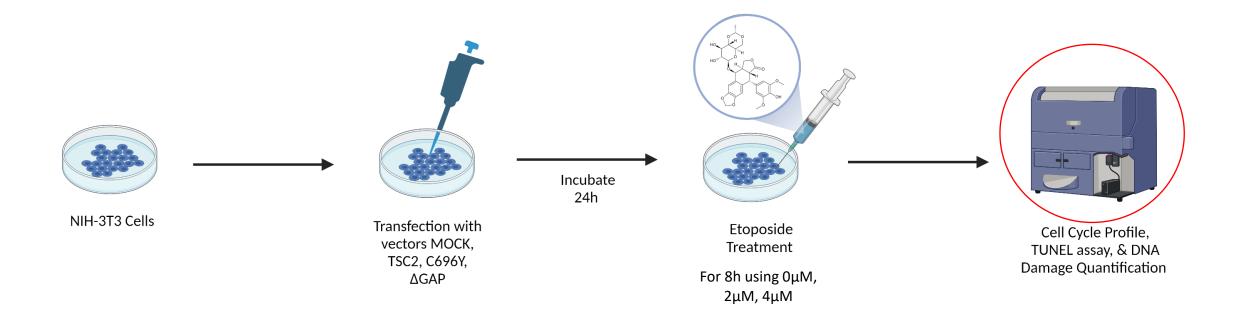
Experimental Design

PORTERLAB



What Etoposide concentration arrests most cells in G2/M?

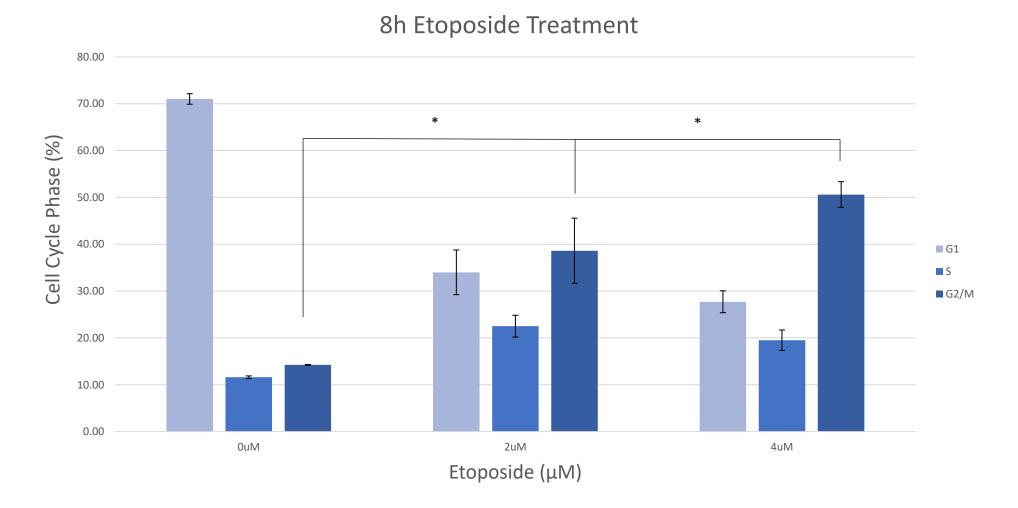
• Objective 1: Etoposide dose curve to obtain G2/M arrest in cell cycle





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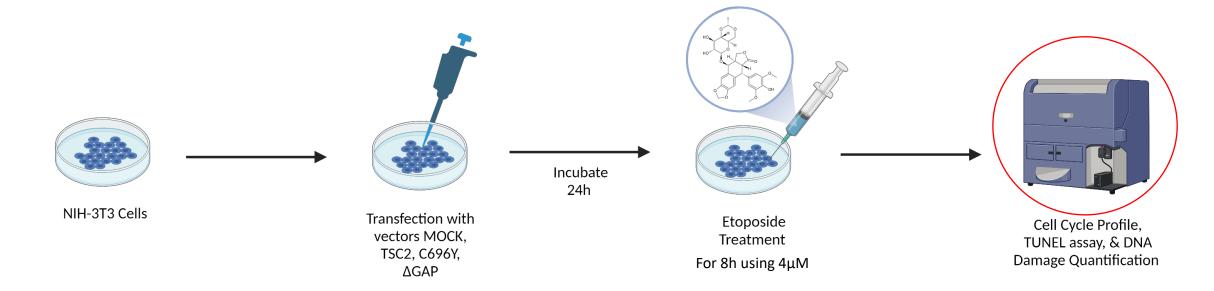
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Percentage of Apoptotic Cells Significantly Greater In TSC2-C696Y Mutant

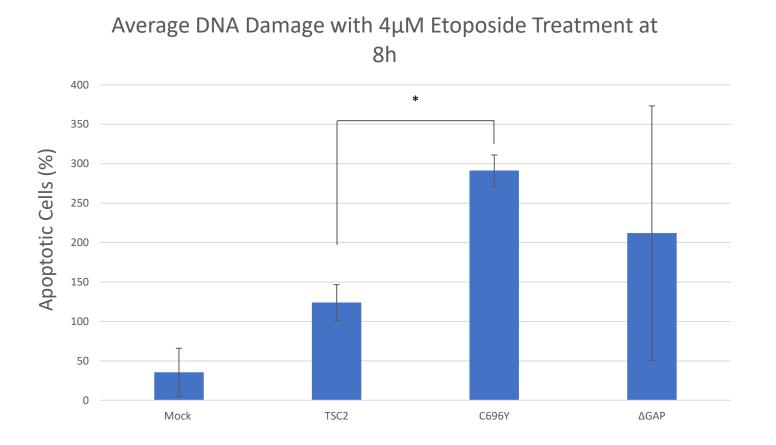
• Objective 2: Analyze apoptotic cell population using TUNEL Assay





Percentage of Apoptotic Cells Significantly Greater In TSC2-C696Y Mutant

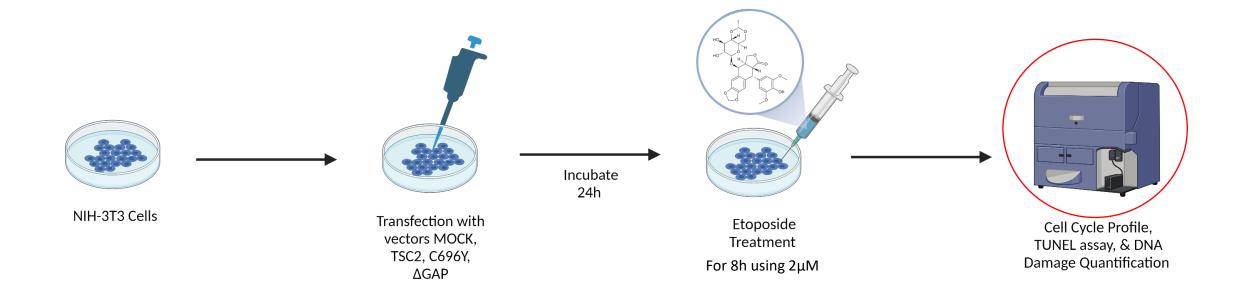
• Objective 2: Analyze apoptotic cell population using TUNEL Assay





DNA Damage Quantification With γH2AX & Flow Cytometry

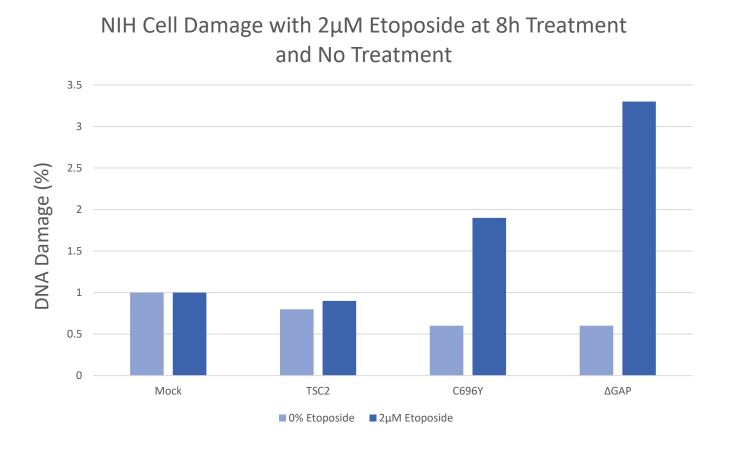
• **Objective 3**: Quantify DNA damage using γH2AX and flow cytometry





DNA Damage Quantification With γH2AX & Flow Cytometry

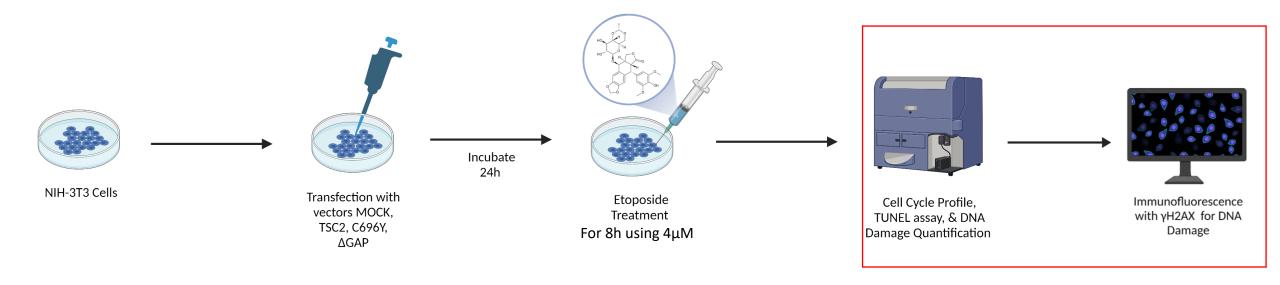
• **Objective 3**: Quantify DNA damage using γH2AX and flow cytometry





DNA Damage Quantification With γH2AX Immunofluorescence

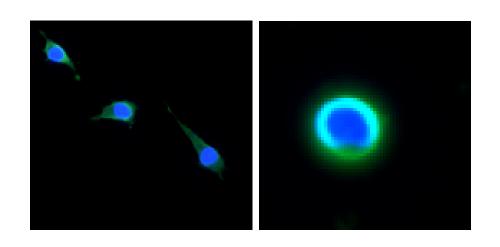
• Objective 3: Quantify DNA damage using γH2AX and Immunofluorescence



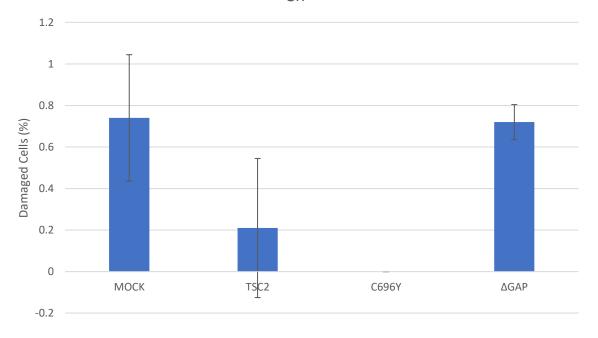


DNA Damage Quantification With γH2AX Immunofluorescence

• Objective 3: Quantify DNA damage using γH2AX and Immunofluorescence

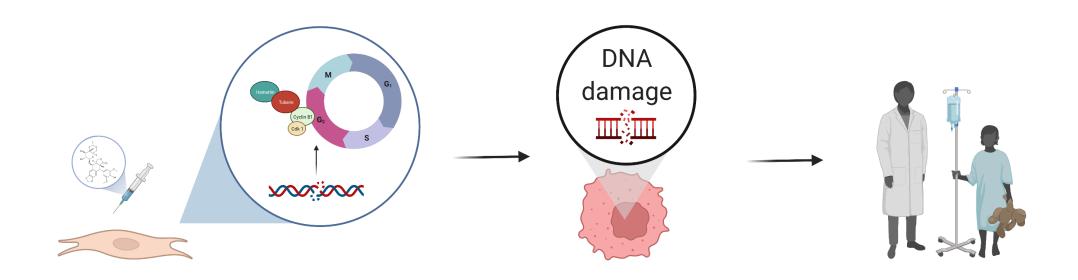


Cells With DNA Damage After $4\mu M$ Etoposide Treatment at 8h





The Importance of Tuberin and DNA Damage



DNA Damage Research

Understanding DNA Damage induced carcinogenesis

Providing better outcomes for patients





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- Supervisors: Dr. Lisa Porter and Dr. Elizabeth Fidalgo da Silva
- Porter Lab Team Members
- Thank you for your support!

