



















PATIENT SAFETY



- Access to new cutting edge treatment, with better outcomes not available outside trial
- First to benefit if treatment has efficacy ( Long term benefits of Immune therapy)
- · Patients monitored very closely for safety
- Cost of treatment covered by study

# Risks



• The new treatment may not be better than, or even as good as standard treatment

· Adverse events may be unexpected, serious









- Expenses extra visits, travel, housing & childcare costs.
- Mandate extra testing- Biopsy, research specimens imaging frequency , uncomfortable & time consuming.
- · Benefits in some patients, may not work for everyone





## Dose Limiting Toxicity (DLT)



- Toxicity that is considered unacceptable (due to severity and/or irreversibility) and limits further dose escalation
- Protocol-specific
- · Typically defined based on toxicity seen in the first cycle
  - Grade 2 intolerable or worse non-hematologic toxicity despite supportive measures
  - · Grade 3 or worse hematologic toxicity
  - Ex Grade 4 Thrombocytopenia

10





#### Maximum tolerated dose (MTD)

• The highest dose of a drug that will be tolerated and does not cause unacceptable side effects

## Optimal biological dose (OBD) - Biomarkers

- Alternative strategy to (MTDs) for identifying the recommended phase II trial doses (RP2Ds).
- The dose that has a desirable efficacy performance while still safeguarding patients with an acceptable toxicity profile





Challenging for patients

- Serial sample collection
- Require admission











## **AESI**

- A noteworthy event for the particular product or class of products that a **sponsor** may wish to **monitor** carefully.
- serious or non-serious (e.g. hair loss, loss of taste, impotence)
- **potential precursors** or **prodromes** for more serious medical conditions in susceptible individuals.







participants







- Protocols -High level of **complexity** and **detail**.
- Trial Oversight Highly regulated and scrutinized
- Data Management (100% source data verification)
- Deviations should be minimal
- Deviations that compromise patient safety need to reported to IRB



