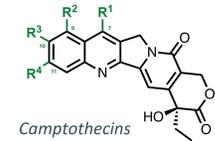


Stuart Barnscher, Raffaele Colombo, Mark Petersen, Michael Brant, Manuel Lasalle, Andrea Hernandez, Renee Duan, Jodi Wong, Samir Das, Vincent Fung, Kaylee Wu, Sam Lawn, Daya Siddappa, Gerry Rowse, and Jamie Rich

Zymeworks Inc. 114 E 4th Ave Suite 800, Vancouver, BC, V5T 1G4, Canada

Background: Camptothecin Therapeutics

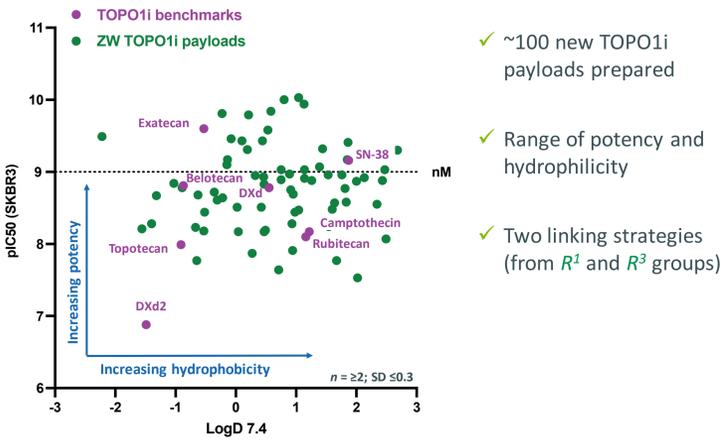


Potent inhibitors of topoisomerase I:

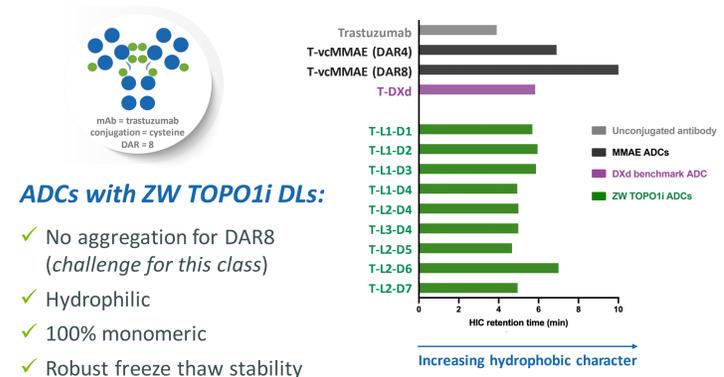
- Discovered in the early 1960 by M. E. Wall and M. C. Wani of Research Triangle Institute
- Isolated from *Camptotheca acuminata* (The Happy Tree)
- Prevents DNA re-ligation which results in double strand breaks and apoptosis

- 3 approved small molecules (Topotecan, Irinotecan, Belotecan)
- 2 approved ADCs (Enhertu®, Trodelvy®)
- Several ADCs, SMDCs, and NPs at different stages of development

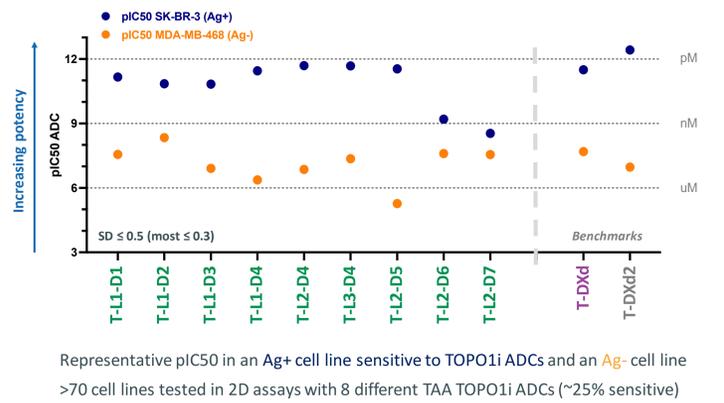
Zymeworks Camptothecin Payloads Span a Range of Potency and Hydrophilicity



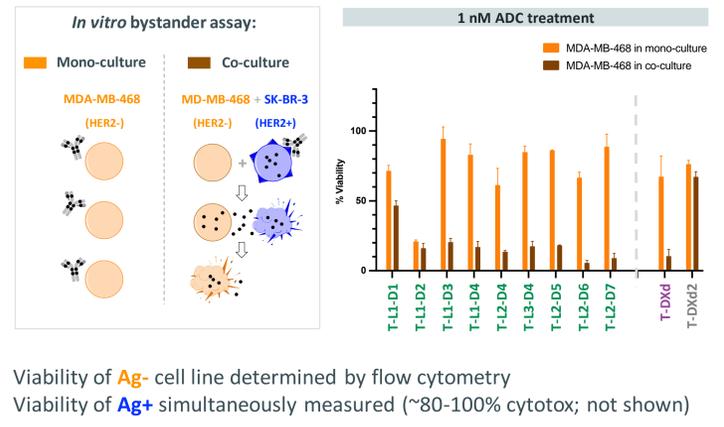
Zymeworks TOPO1i Drug-Linkers Yield ADCs with Favorable Physicochemical Properties and Low Aggregation



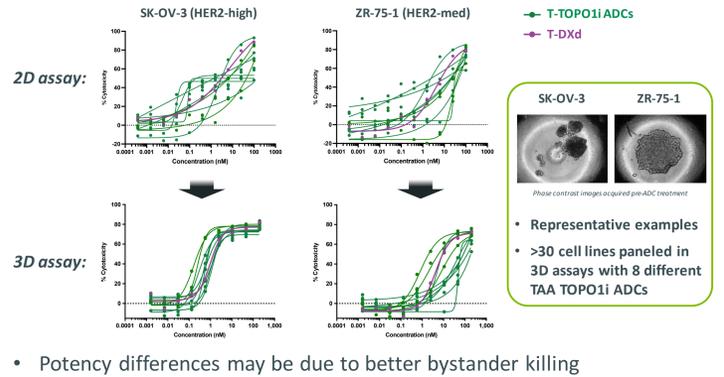
Lead ADCs Showed Good Potency and Selectivity in 2D Cytotoxicity Assays



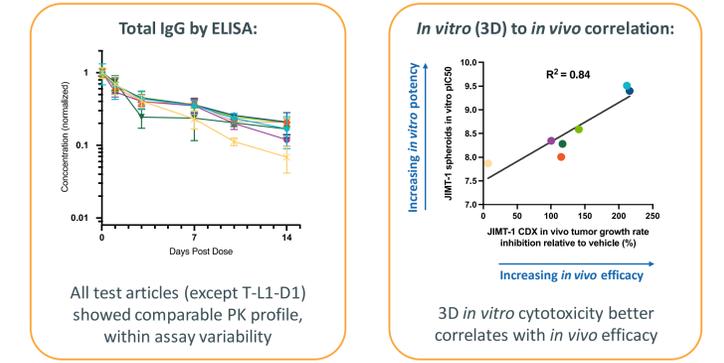
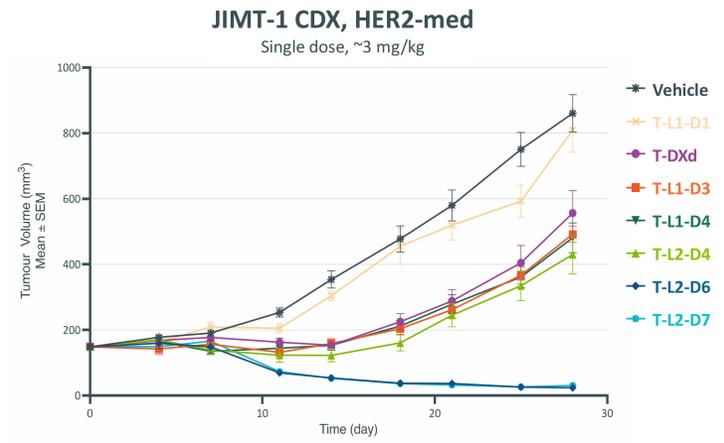
Strong Bystander Activity for Most Zymeworks TOPO1i ADCs



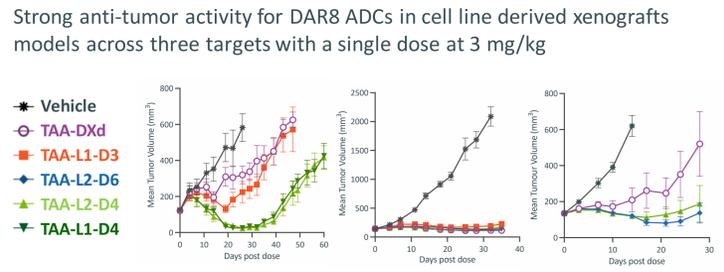
Spheroid Cytotoxicity Assay Altered Dose-Response Relationship and Relative Potency Ranking of ADCs



Zymeworks TOPO1i ADCs Demonstrate Potent in vivo Efficacy

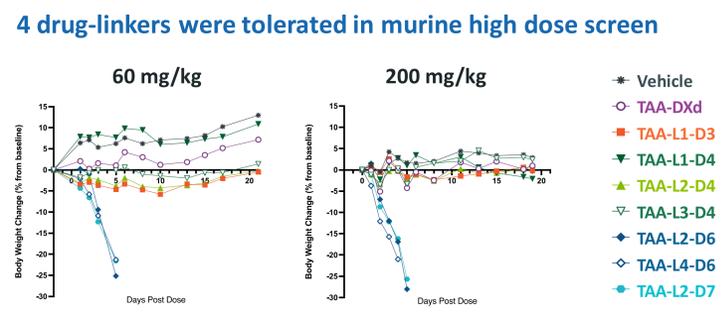


Zymeworks TOPO1i ADCs Demonstrate Anti-tumor Activity in Multiple in vivo Models

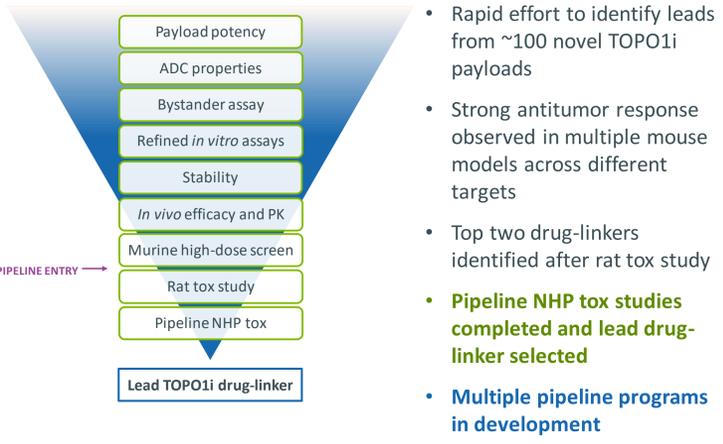


| TAA | TAA1 | TAA2 | TAA3 |
|-------------------------|------------------------|----------|-------------------------|
| Model | Ovarian CDX | Lung CDX | Solid tumor CDX |
| Target expression level | Med/Low, Heterogeneous | High | High/Med, Heterogeneous |
| Mice per group | 6 | 6 | 6 |

Murine and Rat Tolerability Studies Identified 2 Lead Drug Linkers



The Path From Concept to Pipeline



Acknowledgements

Development of this platform would not have been possible without the hard work and effort of the entire ADCDT research group at Zymeworks. Additionally, thank you to the clinical team and senior management for guidance and continued support.