

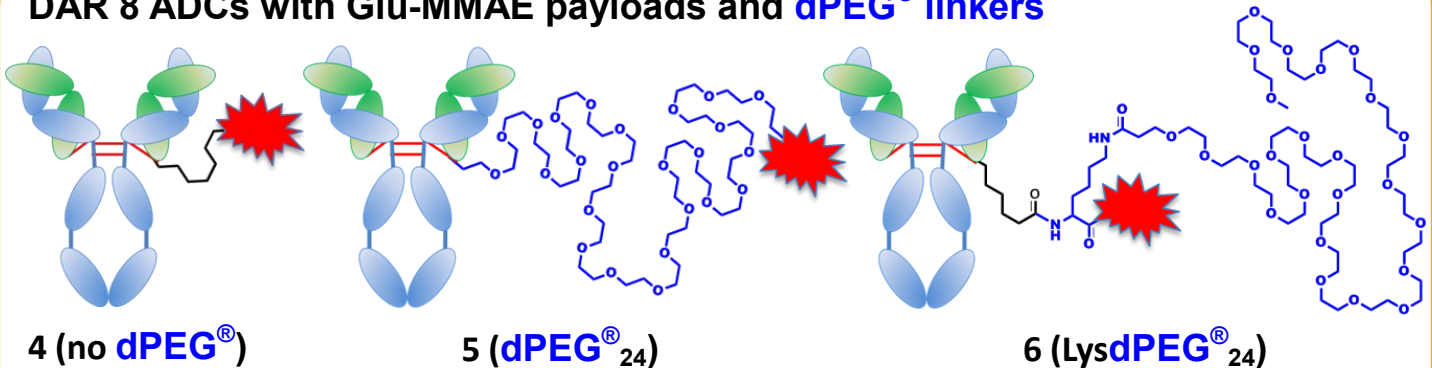


ADCs WITH dPEG® LINKERS :

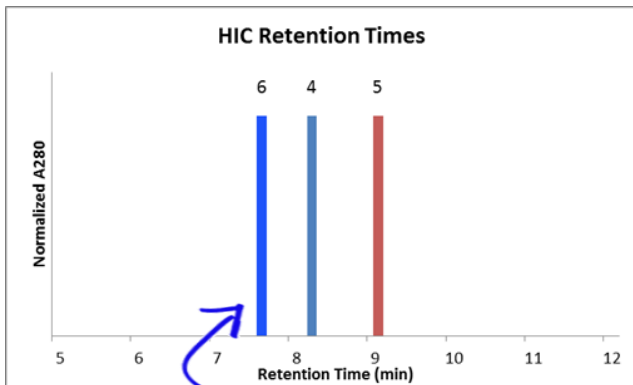
Reducing Hydrophobicity & Clearance

Discrete PEG (dPEG®) linkers are biocompatible structural components that transcend the limitations of traditional linkers, enabling novel conjugate architectures that may display improved *in vivo* performance. For instance, Lyon *et al.* prepared DAR 8 glucuronide-MMAE conjugates with reduced hydrophobicity, reduced clearance, and improved efficacy (*Nat Biotech*, 2015, 33(7), 733-736).

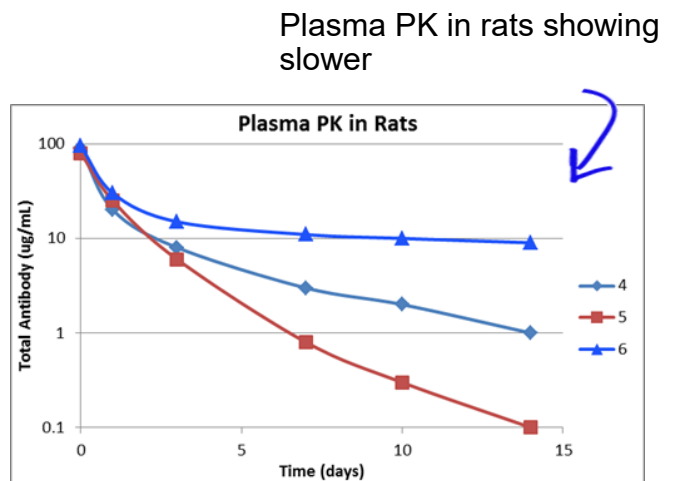
DAR 8 ADCs with Glu-MMAE payloads and dPEG® linkers



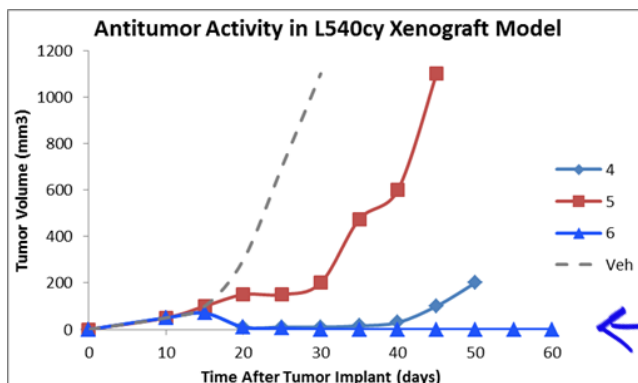
Employing a SuperHydrophilic™ dPEG® linker provides ADC 6 with reduced hydrophobicity, slower clearance, and improved efficacy when compared to other linkers



HIC trace indicating reduced hydrophobicity of 6



Plasma PK in rats showing slower



ADC 6 results in complete tumor regression



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