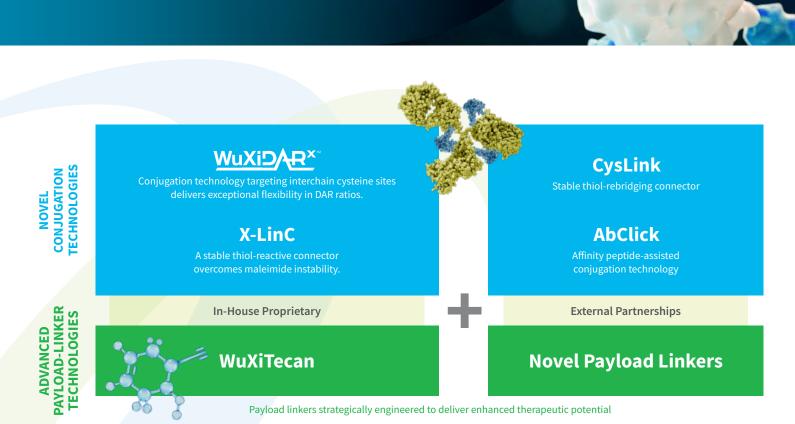


# WuXi XDC TECHNOLOGY PLATFORM

Turn Transformative Potential into Reality

From achieving optimal conjugation stability to ensuring precise payload delivery and minimizing off-target toxicity, every step of bioconjugate development demands cutting-edge solutions. WuXi XDC possesses deep expertise and extensive experience in conjugation. Leveraging our proprietary technologies, we have built a robust foundation for advanced bioconjugate development.

Beyond our internal capabilities, we are actively forging and expanding partnerships with leading industry players. Through these collaborations, we continuously enrich and refine our technology platform, positioning it as the premier destination for comprehensive, customized bioconjugate development solutions.



A Powerful Technology Platform for ADCs and Diverse Bioconjugate Development



Diverse conjugation strategy and payload linker options



Redefine bioconjugate development efficiency and reliability

### **Versatile Drug-to-Antibody Ratio Options with**

### **WuXiD**

### **Proprietary Conjugation Technology**

WuXiDARx, our proprietary conjugation technology targeting the clinically validated interchain cysteine sites, delivers exceptional flexibility in drug-to-antibody ratio (DAR) and demonstrates high homogeneity. It is compatible with native IgG1 antibodies and a wide range of common payload linker and significantly simplifies the CMC process. The technology delivers significant value across the development continuum: When screening antibody and payload-linker combinations during discovery, it streamlines ADC engineering and facilitates optimal DAR identification. At the CMC stage, it accelerates development, mitigates risks, and reduces large-scale manufacturing costs.

### **Features and Advantages**





### **Flexible** DAR

- WuXiDAR1
- WuXiDAR2
- WuXiDAR4 WuXiDAR6



### Highly Homogenous **ADC**

- ≥85% (DAR1, DAR2) or ≥ 65% (DAR4, DAR6) a single conjugated species without column purification
- ≥ 95% with column purification



### Compatible with **Native mAbs**

No antibody engineering modification needed



#### **Meeting Clients' Diversified Needs**

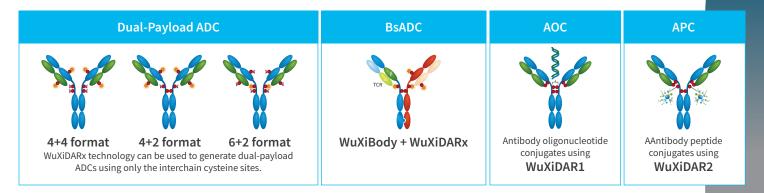
- Compatible with common payload-linkers: MMAE, MMAF, DXD, PBD
- No connector modification



### Simple CMC and **Cost Savings**

- Simple conjugation process
- Comparable COGs with conventional Cys random conjugation

### **Enable Versatility across Diversified Bioconjugates beyond Conventional ADCs**



<sup>\*</sup> WuXiDARx was developed with BioReinno's MCLICK-DAR1-A1, MCLICK-DAR2-A1, and MCLICK-DAR6-A1 technologies.

### **Robust and Validated Conjugation Technology**



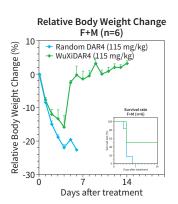
ADCs produced from WuXiDAR4 demonstrated comparable cytotoxicity to ADCs produced by conventional random conjugation methods, but demonstrate better pharmacokinetics (PK) profiles and tolerability. So far, 7 ADC molecules that WuXiDAR4 generated have entered clinical trials.

### **Improved Efficacy and Tolerability**

WuXiDAR4 technology features an enhanced high-homogeneity DAR 4 profile. ADCs engineered with the technology demonstrate superior PK, *in vivo* efficacy, and safety profile relative to ADCs achieving DAR 4 via conventional random cysteine conjugation.

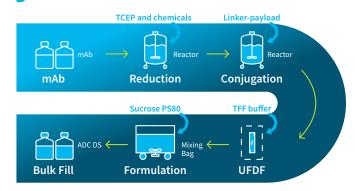
# At dose of 1.25 mg/kg 1500 DPBS Random DAR4 WuXiDAR4 WuXiDAR4 900 7 14 21 28 Days after treatment

WuXiDAR4-ADC shows stronger in vivo efficacy (N87CDX)



WuXiDAR4-ADC shows better tolerability (C57BL/6 mice, single dose)

### Simple and Robust CMC



10+ 7
CMC projects Clinical-stage ADC molecules

2+ kg
GMP batches
prepared

## Ideal for Your Antibody-Oligonucleotide Conjugates



# High Purity Up to ~ 90% D1 species before column purification



#### Simple Process

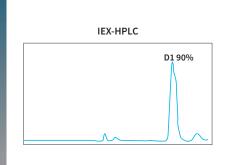
Chemical conjugation, comparable with symmetrical antibody, no sequence engineering or enzymatic reaction required

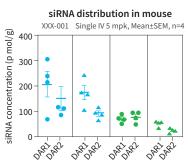


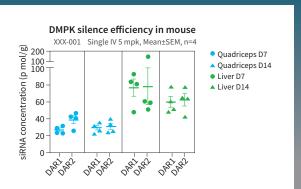


#### **Good Efficacy**

Higher siRNA in target tissue and stronger DMPK silence efficiency in mouse





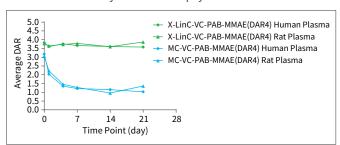


# X-LinC Novel Thiol Reactive Connector

The commonly employed maleimide-thiol conjugation in ADCs is susceptible to retro-Michael addition—a reversible reaction that leads to compromised efficacy and off-target toxicity. WuXi XDC addresses this inherent instability through our proprietary X-LinC connector technology. This innovative solution has demonstrated superior plasma stability, high conjugation efficiency, and seamless compatibility with WuXiDARx.

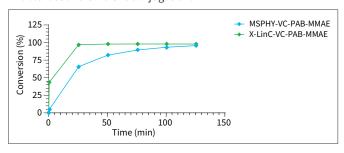
### **Superior stability**

Maintain ADC stability with minimal payload release in circulation.

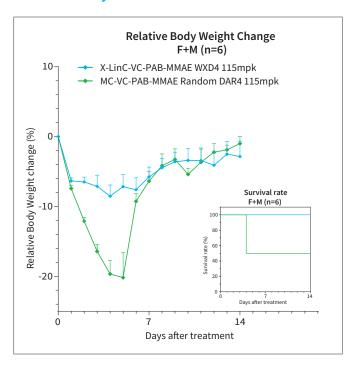


#### **Fast reaction kinetics**

Enable fast and efficient conjugation.



### **Better toxicity tolerance**



# **CysLink**Thiol-Rebridging Connector



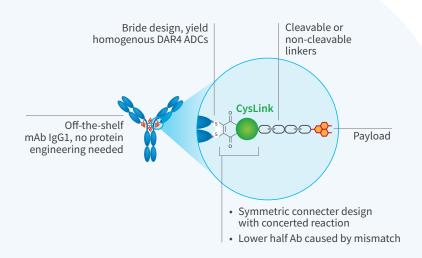
### **Superior stability:**

Forms robust conjugation, ensuring ADC stability and controlled drug release while minimizing toxicity.

### Fully compatible with WuXiDARx

#### Reduced half antibody ratio:

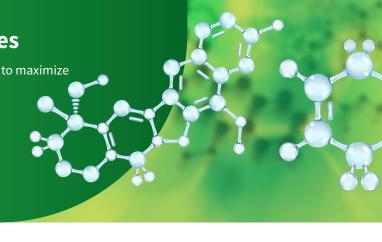
Enhances homogeneity, optimizing conjugation efficiency for more effective therapies.



### **Advanced Payload-Linker Technologies**

Our advanced payload-linker technologies are strategically engineered to maximize therapeutic potential through 3 core innovations:

- o Tunable payload potency balancing efficacy and safety
- Novel cleavage mechanisms enabling controlled drug release
- Hydrophilicity-optimized linkers engineered to enhance safety



### A diverse portfolio of advanced payload-linkers engineered for safety, efficacy, and flexibility across development stages

	Design Strategy		Connector	Release Mechanism	Payload	Stage	
<b>WuXiTecan-1</b> Source: WuXi XDC	Novel TOPi payload with GGFG linker		Mal	Peptidase (Lysosomal)	Novel payload	Preclinical	
	Efficacy	<ul> <li>Payload efficacy is comparable to DXd</li> <li>WuXiTecan-1 ADC efficacy is comparable to DS-8201 in CDX model</li> </ul>					
	Safety	• WuXiTecan-1 ADC is better tolerated than DS-8201 in acute mice toxicity study • WuXiTecan-1 ADC is well tolerated in cyno pre-tox at 55 mpk Q3W*3					

	Design Strategy and Advantages		Connector	Release Mechanism	Payload	Stage	
<b>WuXiTecan-2</b> Source: WuXi XDC	Exatecan with novel hydrophilic linker		Mal	Peptidase (Lysosomal)	Exatecan	Preclinical	
	Hydrophilicity	Better hydrophilicity compared to benchmark*					
	Efficacy	• WuXiTecan-2 ADC shows similar or better efficacy compared to benchmarks* in the CDX model					
	Safety	<ul> <li>WuXiTecan-2 ADC is well tolerated in acute mice toxicity study compared to benchmark*</li> <li>WuXiTecan-2 ADC is well tolerated in cyno pre-tox at 45 mpk Q3W*3</li> </ul>					

 $<sup>{}^{\</sup>star}\mathsf{Benchmark} : \mathsf{Exatecan} \ \mathsf{with} \ \mathsf{hydrophilic} \ \mathsf{linkers} \ \mathsf{that} \ \mathsf{showed} \ \mathsf{promising} \ \mathsf{results} \ \mathsf{in} \ \mathsf{clinical} \ \mathsf{trials}$ 

	Design Strategy and Advantages	Connector	Release Mechanism	Payload	Stage
<b>T-moiety-Exatecan</b> Source: Partner	Exatecan with novel hydrophilic linker	Mal	Peptidase (Lysosomal)	Exatecan	Phase I
OHPAS-Nexatecan Source: Partner	Novel payload and self-immolative adaptor	Mal/IAA	Glucuronidase (Lysosomal)	Nexatecan	СМС
<b>UniLinker-Exatecan</b> Source: Partner	Exatecan with novel TME release linker	Mal	Peptidase (Lysosomal and TME)	Exatecan	СМС

### **About WuXi XDC**

**WuXi XDC** is a leading global CRDMO focused on antibody drug conjugates (ADC) and the broader bioconjugate market. It provides end-to-end contract research, development and manufacturing services for bioconjugates, including ADCs. Its services cover antibody and other bioconjugate intermediates, chemical payloads and linkers, as well as bioconjugate drug substances and drug products.



