Self-Arming Warheads (SAWs) for Antibody-Drug Conjugates (ADC)

**OCR Number:** OCR 6808

**Description:**

- About Self-Arming Warheads (SAWs)
  - SAWs caged as stable, inert Ab cargos inactive until endocytosis
  - Predicted reduced off-target toxicity:
    - Activated SAWs are cell-impermeable
  - Potent (< 100 nM activity)
- Total synthesis at high efficiency
  - Total synthesis facilitates novel chemistry
  - Convergent, high-yielding route to entire framework
  - Few chromatographic purifications
  - Highly amenable to SAR
  - Complete synthetic control
  - Readily conjugated
**Activation via Endocytosis**

- **DNA targeting domain**
- **Prodrug**
- **Degradable linker**

**No Toxic Escape from ADC or Target Cell**

- 100 nM OCR6808-15a SAW causes extensive DNA alkylation
- DNA damage detected using 5 nM of most potent SAW.

**IP Status:** ADC applications unpublished. Provisional patents filed. Both available under CDA.

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