



# **Single Domain Antibody Libraries**

LakePharma has partnered with Twist Biopharma, a division of Twist Bioscience, to offer single domain antibody libraries as part of the antibody discovery and engineering services.

# **Three Single Domain Antibody Libraries**

### **VHH Ratio**

Specific oligo pools model the natural VHH repertoire

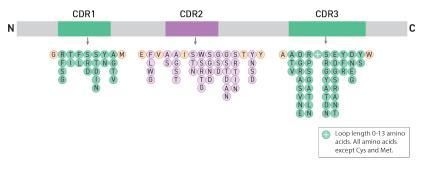
## **VHH Shuffle**

Natural llama CDR sequences in the context of a llama consensus framework

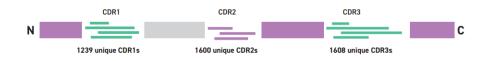
## VHH hShuffle

Natural Ilama CDR sequences in the context of a partially humanized VHH framework

#### **VHH Ratio**



- 2,391 CDR sequences analyzed for position-specific variation
- Controlled CDR diversity introduced in the library
- Consensus Ilama framework



#### **VHH Shuffle**

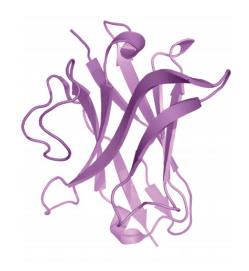
- Each unique CDR individually synthesized
- Shuffled in consensus llama framework
- Final diversity of library greater than theoretical diversity
- Theoretical library diversity of 3.2 x 10<sup>9</sup>

#### VHH hShuffle

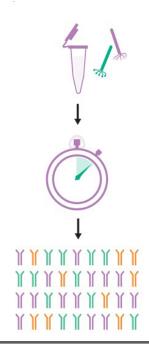
- Shuffled CDRs with theoretical library diversity of 3.2 x 10°
- Partially humanized framework: Framework 1, 3 and 4 were humanized using the human germline DP-47 framework

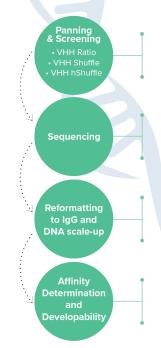
# **Highlights**

- Small & modular antibodies
- Stable & robust
- Easier to engineer & manufacture
- Access to epitopes that are usually sterically hindered by an IgG
- Create building blocks for bispecific antibodies
- Off-the-shelf libraries ready for screening

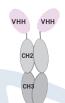


# **Proof of Concept Data**





- 5 successive rounds against protein target
- Phage ELISA screen
- NGS clonal enrichment monitored at each round
- All unique ELISA-positive clones synthesized as VHH-Fc

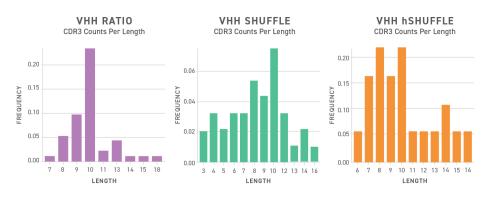


- VHH-Fc affinities determined using Carterra® LSA platform
- VHH-Fc developability assessed using a panel of assays

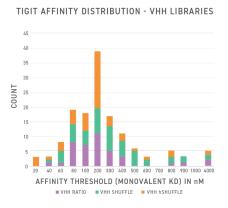
## ELISA + Colony Count: 1 x 384-Well Plate Picked per Library per Round

Library	Round 3	Round 4	Round 5	Uniques
VHH Ratio	58	85	188	47
VHH Shuffle	128	211	287	58
VHH hShuffle	82	222	255	56

# **CDRH3 Length Distributions**



## **Array SPR Binding Analysis**



Anti-TIGIT clones from VHH libraries encompass a range of affinities and diversity

Contact Us

Corporate Headquarters 201 Industrial Road San Carlos, CA 94070 Tel. 650-288-4891

Tel. 888-406-5658 (Toll-free)

Fax 888-635-3618

Email inquiries@lakepharma.com Web www.lakepharma.com