

# B CELL CLONING ON THE BEACON® PLATFORM

## B Cell Cloning

Through a collaboration with Berkeley Lights (BLI), ChemPartner has expanded our antibody discovery service offering to include B Cell cloning. The Beacon® platform, created by BLI, is capable of screening thousands of plasma B cells from immunized animals, which speeds up a traditionally time-consuming hybridoma process.

Plasma B cells secreting antibodies of interest are selected based on cell or bead-based binding and/or functional assays that are run serially or by multiplex format. Individual plasma B cells of interest are selectively exported for antibody sequencing. Using this platform, plasma B cells can be characterized in less than eight hours thus accelerating the antibody discovery process.

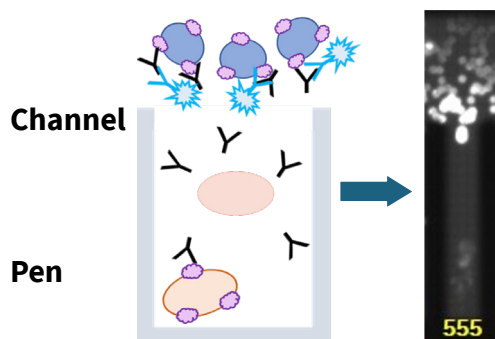


## B Cell Cloning on the Beacon® Platform

- Reduce antibody screening time from 8-12 weeks to one day while interrogating more of the immune repertoire
- Potential to immunize WT or transgenic strains of mice
- Identify plasma B cells from immunized mice with target specific binding and functional activity
- Demonstrated functional blocking of receptor/ligand interactions using PD-L1/PD-1 as test case
- Additional assays such as Ca Flux, cytokine secretion and T cell activation are under development
- Good rate of recovery of single VH/VL sequences from successfully exported plasma B cells

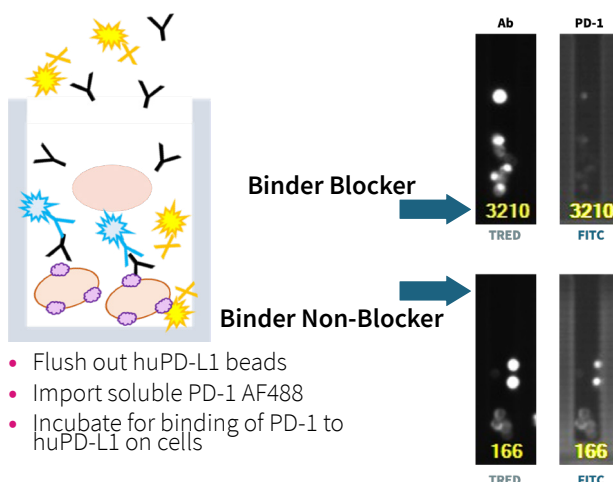
# Assays on Beacon<sup>®</sup>

## In-Channel Bead Binding

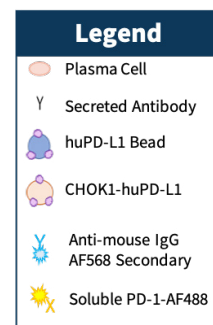


- In-Channel Assay
  - Plasma cell
  - huPD-L1 beads
  - CHOK1-huPD-L1
- Secreted Ab binds to huPD-L1 beads mixed with detection Ab

## In-Pen Binding



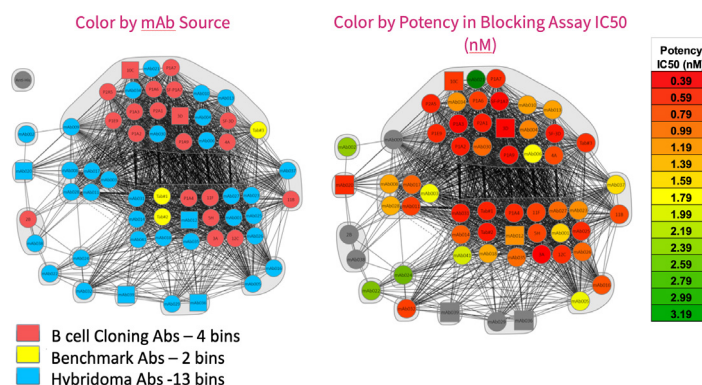
- Flush out huPD-L1 beads
- Import soluble PD-1 AF488
- Incubate for binding of PD-1 to huPD-L1 on cells



## Proof of Concept Study

A POC study was performed using Balb/c mice immunized with recombinant human PD-L1 antigen side by side to compare B cell cloning and hybridoma methods. A majority of PD-L1 blockers identified on the Beacon platform were confirmed in subsequent characterization assays once generated as chimeric IgGs. A majority (58%) of the B cell cloning antibodies had binding and potency similar to or better than benchmark antibodies whereas only 32% of hybridoma antibodies had potency comparable to benchmarks. Thus, B cell cloning enables identification of a greater number of desired hits in a fraction of the time. Antibody candidates with desired properties can be further optimized for manufacturability and developability through ChemPartner's extensive large molecule capabilities.

|   | B Cell Cloning on Beacon | Hybridoma   |
|---|--------------------------|-------------|
| Blockers in Primary Screen                | 35                       | >51         |
| Single Digit nM Affinity                  | 6                        | 12          |
| Blocking False Positive Rate <sup>1</sup> | 5/24 (21%)               | 13/41 (32%) |
| Blocking IC50 ~ Tabs                      | 14/24 (58%)              | 11/41 (32%) |
| Blocking IC50 < Tabs                      | 5/24 (21%)               | 0/41        |
| pM Affinity and Potent Blocking           | 2/2 (100%)               | 1/3 (33%)   |
| Epitope Bins                              | 4                        | 13          |
| T Cell Stimulation EC50 ≤ Tabs            | 9/24 (37.5%)             | 4/41 (9.7%) |



## Typical Output for B Cell Cloning Workflow

