

Next Generation Biologics for Cancer Therapy: Beyond the Success of Conventional Monoclonal Antibodies

Moderator: Mike Rice, Senior Consultant, Defined Health

Panelists:

- Christian Zahnd, PhD, Chief Executive Officer, Molecular Partners AG
- John Haurum, MD, PhD, Chief Executive Officer , F-star
- Hans-Peter Gerber, PhD, Executive Director, Pfizer
- John M. Lambert, Ph.D, Executive Vice President & Chief Scientific Officer, ImmunoGen, Inc.
- Bill Grossman MD, PhD, Senior Vice President of Research & Development, Biothera

The logo features the words "CANCER" and "PROGRESS" in a bold, black, sans-serif font. "CANCER" is positioned above "PROGRESS". Below "PROGRESS" is the tagline "by Defined Health" in a smaller, italicized, black font. The entire text is overlaid on a large, light blue, tilted oval shape that serves as a background for the logo.

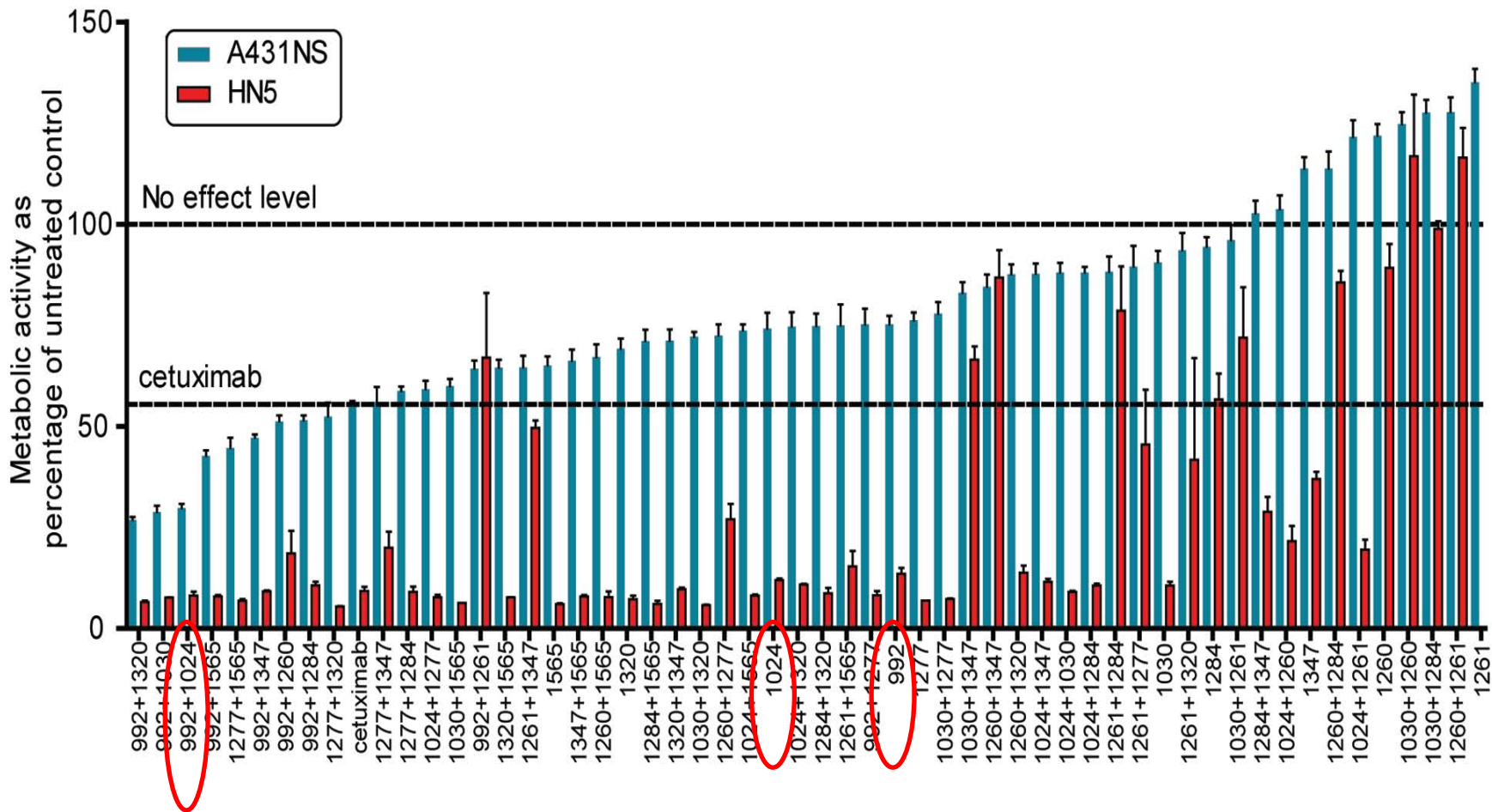
CANCER
PROGRESS
by Defined Health



Advancing Novel Bispecific Antibody Biologics

March 2013

Systematic Screening for Synergistic Combinations



Symphogen: Koefoed et al, mAbs 2011;3:584-95

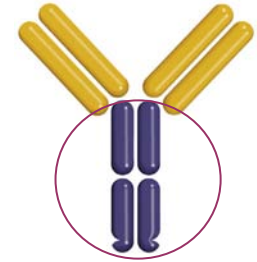
Main Bispecific Issues

- ⌘ Greatest challenge is to find the right combination leading to enhanced efficacy, durability of response, and safety
- ⌘ Not any combination of “best mAb” will work
- ⌘ Druggability issues
 - Manufacturability
 - Stability & PK
- ⌘ The bispecific preferably contributes additional biology not achieved with the combination (efficacy improvement)
 - Synergistic, dual pathway blockade
 - Immune effector function recruitment
 - Tissue targeting of second antibody
- ⌘ Need to focus on biological hypothesis testing



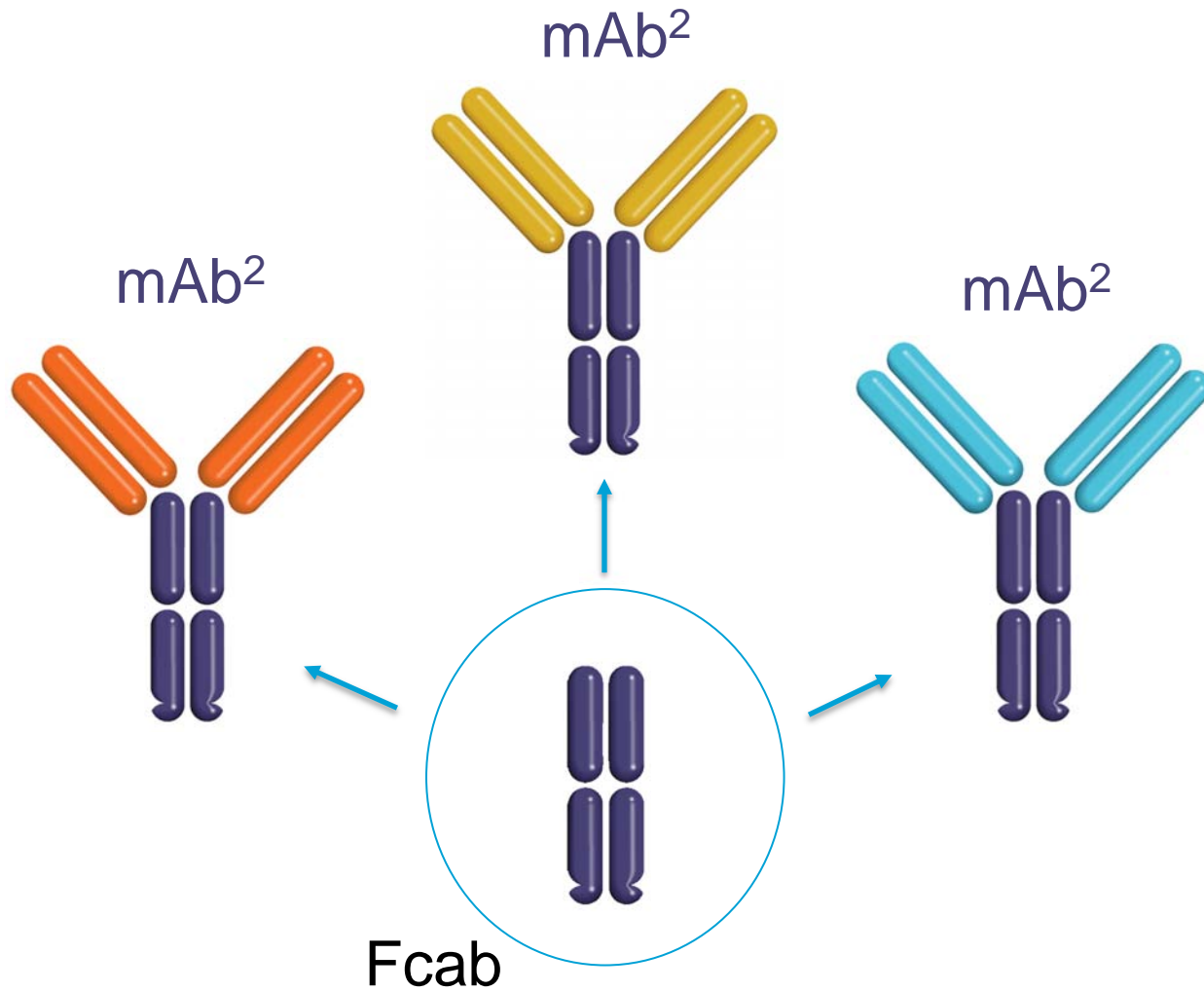
Fcabs: at the heart of Modular Antibody Technology

- ⚡ Drug-like characteristics:
 - Similar potency to a full-length antibody
 - Binds Fc receptors
 - Mobilises immune effector functions



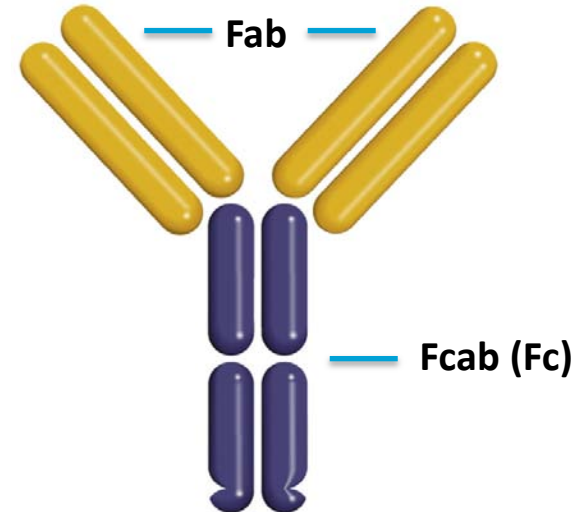
Antigen binding sites

mAb²: speed in unlocking novel biology through modularity



mAb²: A genuine bispecific mAb

- ⌘ Looks like a mAb
 - comparable structure
- ⌘ Works like a mAb
 - retains all antibody functionality
- ⌘ Behaves like a mAb
 - maintains manufacturing characteristics
- ⌘ Performs beyond a mAb
 - synergy through bispecificity
- ⌘ Well-established regulatory path



mAb² rapidly generated from Fcab

Lead Fcab
by Q3 2012

Formatted mAb²
in Q3 2012

In-vivo studies
in Q4 2012

Her2 Fcab

mAb A/Her2 Fcab mAb²

mAb B/Her2 Fcab mAb²

mAb C/Her2 Fcab mAb²

mAb D/Her2 Fcab mAb²

- ⌘ Optimum modality currently being identified
- ⌘ Program to be progressed internally

Summary



- ✦ Powerful modular bispecific antibody platform
- ✦ CMC challenges identical to mAb
- ✦ Lead program funded to IND filing
- ✦ Multiple opportunities for building value through partnerships

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DefinedHealth
unconventional insight



F-star