1971 ELISA assay developed independently by Eva Engvall and Peter Perlman (21)

1972 FACS instrument developed and patented by Len Herzenberg's lab at Stanford University (22)

1975 Georges Köhler and César Milstei develop hybridomas leading to the production of mAbs (23)

1976 Susumu Tonegawa describes somatic recombination of immunoglobulin genes to account for incredible diversity (24)

> 1978 Hybritech becomes the first mAb company (25)

1979 First patent on hybridoma technology awarded to Wistar Institute (26) Western blotting, perhaps the most widely used immunoassay in research, is invented by Harry Towbin et al. (27)

1980

990

1985 Immunoprecipitation (ChIP) assay (32)

Sean P. O'Neill and Joseph Wu awarded patent for quantitative immunoprecipitation assay (33)

1990 John McCafferty et al. report the

use of phage display for antibody discovery (34)

995

Katherine Knight and colleagues at Loyola University, Chicago, USA published first paper on rabbit mAb development (35)

1996 Prostascint®, radio-labeled anti-

PSMA (prostate specific membrane antigen) imaging antibody approved by the FDA (36)

1997 Idec markets the world's

first mAb treatment for lymphoma (Rituxan®) (37) 1008

1960

Rosalyn Yalow and Solomon Berson (18

1960

1981 The lab of Herman Eisen develops the first anti-pTyr antibody (28) Hybritech delivers first mAb product to measure IgE in blood to diagnose allergic reactions (29)

1970

1982 Angus Nairn, et al. develop the first phospho-specific antibodies (30)

1984 hCG antibodies used to develop 5 minute pregnancy test (31)

John Lis and David Gilmour develop Chromatin

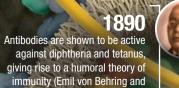
1986

125 YEARS

Antibody Discovery and Development

From early explorations of vaccination to present-day clinical trials, antibody-based research and therapies have long demonstrated their enormous potential to benefit human health.

This year we celebrate the 125th anniversary of the use of antibodies to treat diphtheria and tetanus and the advancement of the humoral theory of immunity. We encourage you to explore the rich history of antibodies and to share our passion for research supporting the next wave of innovations in this field.



ato Shibasaburo) (1)

1965 Thomas Tomasi identifies secretory immunoglobulins (IgA) (19) First fluorescence based assay developed by Martin Fulwyler (20)

1966 Kimishige Ishizaka et al. and S.G.O. Johansson & Hans Bennich independently identified IgE as the reaginic antibody (reviewed in 7)

1967 Kimishige Ishizaka identifies IgE as the reaginic antibody, binding the molecule that induced its synthesis (reviewed in 7)



1950

1953 Wallace Coulter awarded a patent on Coulter principle, enabling flow cytometry (12)

> 1955 Niels Jerne proposes natural-selection theory of antibody formation (13)

1956

Kappa and lambda light chains, then known as Bence Jones proteins, are shown to be two separate proteins by Leonard Korngold and Rose Lipari (reviewed in 7)

1957 Clonal selection theory proposed by Frank

MacFarlane Burnet and David W. Talmage (14,15)



1959 and 1962

Different regions of antibody structures independently elucidated by Gerald Edelman and Rodney Porter (16,17)



1940

Karl Landsteiner and Alexander Wein identify Rh antigens (8)

1942

Albert Coons labels antibodies with FITC orig nating the field of immunofluorescence (9) Jules Freund and Katherine McDermott demonstrate use of adjuvants to stimulate antibody production (10)

1944

IgM is described independently by Jan Waldenström with Kai Pedersen as well as Henry Kunkel (reviewed in 7)

1948 Astrid Fagreaus discovered antibody oduction in plasma B cells (11)

\bigcirc 1930

1938 John Marrack proposes Antigen Antibody binding hypothesis (6)

1939 Arne Tiselius and Elvin Kabat discover the first tibody isotype, gamma-globulin (reviewed in 7)

CST releases The CST Guide: Pathways & Protocols,

a comprehensive scientific and technical resource

for cell biology researchers (46)

¥ 2015

() 2010

Herceptin[®] approved for breas cancer treatment (38)

¥

1999 CST established as an independent company and releases its first substrat motif antibody (#9611) (39)

-0

2000

1891 ve transferrable immunity I von Behring and Kitasato Shibasaburo) (2)

1896 rial, heat-labile serum component (3)

1909

1900

theory (4)

Paul Erlich develops tibody formation **9** 1900

Amroth Wright publishes, Studies on Immunisation", a collection of papers describing opsonization in the context of therapeutic immunization (5)

2000 Abgenix develops XenoMouse[®] which produces fully human antibodies (40)

2004 Erbitux[®] approved by FDA for reatment of colorectal cancer (41)

2006 CST released its first XP® antibody developed using the proprietary XMT[®] method (42)

> 2012 CST publishes NG-XMT[™] method, a proteomics ¥ ach to developing mAbs (43)

2013 Kadcyla® (ado-trastuzumab emtansine), an antibody-drug conjugate, receives FDA approval as late-stage breast cancer treatment (44)

2014 Yervoy[®] (ipilimumab), a monoclonal anti-CTLA4 antibody and the first immune checkpoint cancer therapy receives FDA approval as a latestage melanoma treatment (45)

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www.cellsignal.com/125history

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