

---

**6. REFERENCES**

- Alberola Ila, J., Takaki, S., Kerner, J.D., and Perlmutter, R.M. (1997).** Differential signaling by lymphocyte antigen receptors. *Annu. Rev. Immunol.* 15, 125-154.
- Alderson, M.R., Tough, T.W., Davis Smith, T., Braddy, S., Falk, B., Schooley, K.A., Goodwin, R.G., Smith, C.A., Ramsdell, F., and Lynch, D.H. (1995).** Fas ligand mediates activation-induced cell death in human T lymphocytes. *J. Exp. Med.* 181, 71-77.
- Amakawa, R., Hakem, A., Kundig, T.M., Matsuyama, T., Simard, J.J., Timms, E., Wakeham, A., Mittruecker, H.W., Griesser, H., Takimoto, H., Schmits, R., Shahinian, A., Ohashi, P., Penninger, J.M., and Mak, T.W. (1996).** Impaired negative selection of T cells in Hodgkin's disease antigen CD30-deficient mice. *Cell* 84, 551-562.
- Amos, L.A. (1991).** Molecules of the cytoskeleton. New York: Guilford Press.
- Aruffo, A., Stamenkovic, I., Melnick, M., Underhill, C.B., and Seed, B. (1990).** CD44 is the principal cell surface receptor for hyaluronate. *Cell* 61, 1303-1313.
- Babbitt, B.P., Allen, P.M., Matsueda, G., Haber, E., and Unanue, E.R. (1985).** Binding of immunogenic peptides to Ia histocompatibility molecules. *Nature* 317, 359-361.
- Bachmann, M., Barner, M., and Kopf, M. (1999).** CD2 sets quantitative thresholds in T cell activation. *J. Exp. Med.* 190, 1383-1391.
- Bagrodia, S., Derijard, B., Davis, R.J., and Cerione, R.A. (1995).** Cdc42 and PAK-mediated signaling leads to Jun kinase and p38 mitogen-activated protein kinase activation. *J. Biol. Chem.* 270, 27995-27998.
- Barber, E.K., Dasgupta, J.D., Schlossman, S.F., Trevillyan, J.M., and Rudd, C.E. (1989).** The CD4 and CD8 antigens are coupled to a protein-tyrosine kinase (p56lck) that phosphorylates the CD3 complex. *Proc. Natl. Acad. Sci. U. S. A.* 86, 3277-3281.
- Beggs, H.E., Soriano, P., and Maness, P.F. (1994).** NCAM-dependent neurite outgrowth is inhibited in neurons from Fyn-minus mice. *J. Cell Biol.* 127, 825-833.
- Berg, E.L., Goldstein, L.A., Jutila, M.A., Nakache, M., Picker, L.J., Streeter, P.R., Wu, N.W., Zhou, D., and Butcher, E.C. (1989).** Homing receptors and vascular addressins: cell adhesion molecules that direct lymphocyte traffic. *Immunol. Rev.* 108, 5-18.
- Berg, N.N. and Ostergaard, H.L. (1995).** Characterization of intercellular adhesion molecule-1 (ICAM-1)-augmented degranulation by cytotoxic T cells. ICAM-1 and anti-CD3 must be co-localized for optimal adhesion and stimulation. *J. Immunol.* 155, 1694-1702.

- Berg, N.N., Puente, L.G., Dawicki, W., and Ostergaard, H.L. (1998).** Sustained TCR signaling is required for mitogen-activated protein kinase activation and degranulation by cytotoxic T lymphocytes. *J. Immunol.* 161, 2919-2924.
- Bluestone, J. A. (1995).** New perspectives of CD28-B7-mediated T cell costimulation. *Immunity* 2:555-559.
- Borland, G., Ross, J.A., and Guy, K. (1998).** Forms and functions of CD44. *Immunology*. 93, 139-148.
- Bourguignon, L.Y., Lokeshwar, V.B., He, J., Chen, X., and Bourguignon, G.J. (1992).** A CD44-like endothelial cell transmembrane glycoprotein (GP116) interacts with extracellular matrix and ankyrin. *Mol. Cell Biol.* 12, 4464-4471.
- Bourguignon, L.Y., Lokeshwar, V.B., Chen, X., and Kerrick, W.G. (1993).** Hyaluronic acid-induced lymphocyte signal transduction and HA receptor (GP85/CD44)-cytoskeleton interaction. *J. Immunol.* 151, 6634-6644.
- Bourguignon, L.Y. and Jin, H. (1995).** Identification of the ankyrin-binding domain of the mouse T-lymphoma cell inositol 1,4,5-trisphosphate (IP3) receptor and its role in the regulation of IP3-mediated internal Ca<sup>2+</sup> release. *J. Biol. Chem.* 270, 7257-7260.
- Bourguignon, L.Y., Gunja Smith, Z., Iida, N., Zhu, H.B., Young, L.J., Muller, W.J., and Cardiff, R.D. (1998).** CD44v(3,8-10) is involved in cytoskeleton-mediated tumor cell migration and matrix metalloproteinase (MMP-9) association in metastatic breast cancer cells. *J. Cell Physiol.* 176, 206-215.
- Brdicka, T., Cerny, J., and Horejsi, V. (1998).** T cell receptor signalling results in rapid tyrosine phosphorylation of the linker protein LAT present in detergent-resistant membrane microdomains. *Biochem. Biophys. Res. Commun.* 248, 356-360.
- Brown, D. (1993).** The tyrosine kinase connection: how GPI-anchored proteins activate T cells. *Curr. Opin. Immunol.* 5, 349-354.
- Brown, D.A. and London, E. (1997).** Structure of detergent-resistant membrane domains: does phase separation occur in biological membranes? *Biochem. Biophys. Res. Commun.* 240, 1-7.
- Brown, J.L., Stowers, L., Baer, M., Trejo, J., Coughlin, S., and Chant, J. (1996).** Human Ste20 homologue hPAK1 links GTPases to the JNK MAP kinase pathway. *Curr. Biol.* 6, 598-605.
- Brown, S.S. (1993).** Phenotypes of cytoskeletal mutants. *Curr. Opin. Cell Biol.* 5, 129-134.
- Brown, T.A., Bouchard, T., St.John, T., Wayner, E., and Carter, W.G. (1991).** Human keratinocytes express a new CD44 core protein (CD44E) as a heparan-sulfate intrinsic membrane proteoglycan with additional exons. *J. Cell Biol.* 113, 207-221.

- Brunner, T., R. Mogil, D. LaFace, N. J. Yoo, A. Mahboubi, F. Echeverri, S. j. Marin, W. R. Force, D. H. Lynch, C. F. Ware, and D. R. Green. (1995).** Cell-autonomous Fas (CD95)/Fas-ligand interaction mediated activation-induced apoptosis in T cell hybridomas. *Nature* 373:441-444.
- Bubeck Wardenburg, J., Fu, C., Jackman, J.K., Flotow, H., Wilkinson, S.E., Williams, D.H., Johnson, R., Kong, G., Chan, A.C., and Findell, P.R. (1996).** Phosphorylation of SLP-76 by the ZAP-70 protein-tyrosine kinase is required for T-cell receptor function. *J. Biol. Chem.* 271, 19641-19644.
- Bubeck Wardenburg, J., Pappu, R., Bu, J.Y., Mayer, B., Chernoff, J., Straus, D., and Chan, A.C. (1998).** Regulation of PAK activation and the T cell cytoskeleton by the linker protein SLP-76. *Immunity*. 9, 607-616.
- Cantrell, D. (1996).** T cell antigen receptor signal transduction pathways. *Annu. Rev. Immunol.* 14:259-274.
- Caplan, S. and Baniyash, M. (1995).** Multisubunit receptors in the immune system and their association with the cytoskeleton: in search of functional significance. *Immunol. Res.* 14, 98-118.
- Chambers, C.A. and Allison, J.P. (1997).** Co-stimulation in T cell responses. *Curr. Opin. Immunol.* 9, 396-404.
- Chan, A.C., Iwashima, M., Turck, C.W., and Weiss, A. (1992).** ZAP-70: a 70 kd protein-tyrosine kinase that associates with the TCR zeta chain. *Cell* 71, 649-662.
- Chan, A. C., and A. S. Shaw, (1996).** Regulation of antigen receptor signal transduction by protein tyrosine kinases. *Curr. Opin. Immunol.* 8, 394-401.
- Collins, T.L., Deckert, M., and Altman, A. (1997).** Views on Vav. *Immunol. Today*. 18, 221-225.
- Colombatti, A., Hughes, E.N., Taylor, B.A., and August, J.T. (1982).** Gene for a major cell surface glycoprotein of mouse macrophages and other phagocytic cells is on chromosome 2. *Proc. Natl. Acad. Sci. U. S. A.* 79, 1926-1929.
- Conrad, P., Rothman, B.L., Kelley, K.A., and Blue, M.L. (1992).** Mechanism of peripheral T cell activation by coengagement of CD44 and CD2. *J. Immunol.* 149, 1833-1839.
- Cooper, D.L., Dougherty, G., Harn, H.J., Jackson, S., Baptist, E.W., Byers, J., Datta, A., Phillips, G., and Isola, N.R. (1992).** The complex CD44 transcriptional unit; alternative splicing of three internal exons generates the epithelial form of CD44. *Biochem. Biophys. Res. Commun.* 182, 569-578.

- Crabtree, G.R. (1989).** Contingent genetic regulatory events in T lymphocyte activation. Science 243, 355-361.
- Crespo, P., Schuebel, K.E., Ostrom, A.A., Gutkind, J.S., and Bustelo, X.R. (1997).** Phosphotyrosine-dependent activation of Rac-1 GDP/GTP exchange by the vav proto-oncogene product. Nature 385, 169-172.
- Croft, M. and Dubey, C. (1997).** Accessory molecule and costimulation requirements for CD4 T cell response. Crit. Rev. Immunol. 17, 89-118.
- D'Ambrosio, D., Cantrell, D.A., Frati, L., Santoni, A., and Testi, R. (1994).** Involvement of p21ras activation in T cell CD69 expression. Eur. J. Immunol. 24, 616-620.
- Darnell J., H. Lodish, and D. Baltimore. (1986).** Molecular Cell Biology. pp 771-815. Scientific American Books, New York.
- Davis, M.M., Boniface, J.J., Reich, Z., Lyons, D., Hampl, J., Arden, B., and Chien, Y. (1998).** Ligand recognition by alpha beta T cell receptors. Annu. Rev. Immunol. 16, 523-544.
- Davis, S.J., and van der Merwe, P.A. (1996).** The structure and ligand interactions of CD2: implications for T cell function. Immunol. Today 17, 177-187.
- DeGrendele, H.C., Estess, P., Picker, L.J., and Siegelman, M.H. (1996).** CD44 and its ligand hyaluronate mediate rolling under physiologic flow: a novel lymphocyte-endothelial cell primary adhesion pathway. J. Exp. Med. 183, 1119-1130.
- DeGrendele, H.C., Estess, P., and Siegelman, M.H. (1997).** Requirement for CD44 in activated T cell extravasation into an inflammatory site. Science 278, 672-675.
- Denning, S.M., Le, P.T., Singer, K.H., and Haynes, B.F. (1990).** Antibodies against the CD44 p80, lymphocyte homing receptor molecule augment human peripheral blood T cell activation. J. Immunol. 144, 7-15.
- De Petris, S. (1974).** Inhibition and reversal of capping by cytochalasin B, vinblastine and colchicine. Nature 250:54-56.
- Dhein, J., H. C. Walczac, K. M. Baumler, Debatin, and P. H. Krammer. (1995).** Autocrine T cell suicide mediated by APO-1/(Fas/CD95). Nature 373:438-441.
- Dianzani, U., Bragardo, M., Tosti, A., Ruggeri, L., Volpi, I., Casucci, M., Bottarel, F., Feito, M.J., Bonissoni, S., and Velardi, A. (1999).** CD44 signaling through p56lck involves lateral association with CD4 in human CD4+ T cells. Int. Immunol. 11, 1085-1092.
- Downward, J., Riehl, R., Wu, L., and Weinberg, R.A. (1990).** Identification of a nucleotide exchange-promoting activity for p21ras. Proc. Natl. Acad. Sci. U. S. A. 87, 5998-6002.

- Draberova, L., Amoui, M., and Draber, P. (1996).** Thy-1-mediated activation of rat mast cells: the role of Thy-1 membrane microdomains. *Immunology.* 87, 141-148.
- D'Souza Schorey, C., Boettner, B., and Van Aelst, L. (1998).** Rac regulates integrin-mediated spreading and increased adhesion of T lymphocytes. *Mol. Cell Biol.* 18, 3936-3946.
- Dustin, M.L., M. W. Olszowy, A. D. Holdorf, J. Li, S. Bromley, N. Desai, P. Widder, F. Rosenberger, P. A. van der Merwe, P. M. Allen, and A. S. Shaw. (1998).** A novel adaptor protein orchestrates receptor patterning and cytoskeletal polarity in T-cell contacts. *Cell* 94:667-677.
- Dustin, M.L. and Springer, T.A. (1989).** T-cell receptor crosslinking transiently stimulates adhesiveness through LFA-1. *Nature* 341, 619-624.
- Engvall, E. and Perlman, P. (1971).** Enzyme-linked immunosorbent assay (ELISA). Quantitative assay of immunoglobulin G. *Immunochemistry.* 8, 871-874.
- Ferrell, J.E.J. (1996).** Tripping the switch fantastic: how a protein kinase cascade can convert graded inputs into switch-like outputs. *Trends. Biol. Sci.* 21, 460-466.
- Finco, T.S., Kadlecak, T., Zhang, W., Samelson, L.E., and Weiss, A. (1998).** LAT is required for TCR-mediated activation of PLCgamma1 and the Ras pathway. *Immunity.* 9, 617-626.
- Fischer, K.D., Y. Y. Kong, H. Nishina, K. Tedford, L. E. Marengere, I. Kozieradzki, T. Sasaki, M. Starr, G. Chan, S. Gardener, M. P. Nghiem, D. Bouchard, M. Barbacid, A. Bernstein, and J. M. Penninger. (1998).** Vav is a regulator of cytoskeletal reorganization mediated by the T- cell receptor. *Curr. Biol.* 8:554-562.
- Franklin, R.A., Tordai, A., Patel, H., Gardner, A.M., Johnson, G.L., and Gelfand, E.W. (1994).** Ligation of the T cell receptor complex results in activation of the Ras/Raf-1/MEK/MAPK cascade in human T lymphocytes. *J. Clin. Invest.* 93, 2134-2140.
- Friedrichson, T. and Kurzchalia, T.V. (1998).** Microdomains of GPI-anchored proteins in living cells revealed by crosslinking. *Nature* 394, 802-805.
- Frogner, F.J. and O'Neill, H.C. (1992).** Lymphocyte recirculation: the need for site-specific receptors to dictate T-lymphocyte localization into different tissue sites [editorial]. *Scand. J. Immunol.* 35, 627-632.
- Galandrini, R., Albi, N., Tripodi, G., Zarcone, D., Terenzi, A., Moretta, A., Grossi, C.E., and Velardi, A. (1993).** Antibodies to CD44 trigger effector functions of human T cell clones. *J. Immunol.* 150, 4225-4235.
- Galandrini, R., De Maria, R., Piccoli, M., Frati, L., and Santoni, A. (1994).** CD44 triggering enhances human NK cell cytotoxic functions. *J. Immunol.* 153, 4399-4407.

- Galluzzo, E., Albi, N., Fiorucci, S., Merigliola, C., Ruggeri, L., Tosti, A., Grossi, C.E., and Velardi, A. (1995).** Involvement of CD44 variant isoforms in hyaluronate adhesion by human activated T cells. *Eur. J. Immunol.* 25, 2932-2939.
- Garcia, K. C., C. A. Scott, A. Brunmark, F. R. Carbone, P. A. Peterson, I. A. Wilson, and L. Teyton. (1996).** CD8 enhances formation of stable T-cell receptor/MHC class I molecule complexes [see comments] [published erratum appears in *Nature* 1997 Jun 5;387(6633):634]. *Nature* 384:577-581.
- Gauen, L. K., Y. Zhu, F. Letourneur, Q. Hu, J. B. Bolen, L. A. Matis, R. D. Klausner, and A. S. Shaw. (1994).** Interactions of p59fyn and ZAP-70 with T-cell receptor activation motifs: defining the nature of a signalling motif. *Mol. Cell Biol.* 14:3729-3741.
- Geiger, B., D. Rosen, and G. Berke. (1982).** Spatial relationships of microtubule-organizing centers and the contact area of cytotoxic T lymphocytes and target cells. *J Cell Biol.* 95:137-143.
- Geppert, T.D. and Lipsky, P.E. (1991).** Association of various T cell-surface molecules with the cytoskeleton. Effect of cross-linking and activation. *J. Immunol.* 146, 3298-3305.
- Gogolak, P., Rethy, B., Horvath, A., Toth, G.K., Cervenak, L., Laszlo, G., and Rajnavolgyi, E. (1996).** Collaboration of TCR-, CD4- and CD28-mediated signalling in antigen-specific MHC class II-restricted T-cells. *Immunol. Lett.* 54, 135-144.
- Goodfellow, P.N., Banting, G., Wiles, M.V., Tunnacliffe, A., Parkar, M., Solomon, E., Dalchau, R., and Fabre, J.W. (1982).** The gene, MIC4, which controls expression of the antigen defined by monoclonal antibody F10.44.2, is on human chromosome 11. *Eur. J. Immunol.* 12, 659-663.
- Grakoui, A., S. K. Bromley, C. Sumen, M. M. Davis, A. S. Shaw, P. M. Allen, and M. L. Dustin. (1999).** The immunological synapse: A molecular machine controlling T cell activation. *Science* 285:221-227
- Green, J.M. and Thompson, C.B. (1994).** Modulation of T cell proliferative response by accessory cell interactions. *Immunol. Res.* 13, 234-243.
- Guo, Y., Wu, Y., Shinde, S., Sy, M.S., Aruffo, A., and Liu, Y. (1996).** Identification of a costimulatory molecule rapidly induced by CD40L as CD44H. *J. Exp. Med.* 184, 955-961.
- Guo, Y.J., Ma, J., Wong, J.H., Lin, S.C., Chang, H.C., Bigby, M., and Sy, M.S. (1993).** Monoclonal anti-CD44 antibody acts in synergy with anti-CD2 but inhibits anti-CD3 or T cell receptor-mediated signaling in murine T cell hybridomas. *Cell Immunol.* 152, 186-199.
- Guo, Y.J., Lin, S.C., Wang, J.H., Bigby, M., and Sy, M.S. (1994).** Palmitoylation of CD44 interferes with CD3-mediated signaling in human T lymphocytes. *Int. Immunol.* 6, 213-221.

- Hall, A. (1998).** G proteins and small GTPases: distant relatives keep in touch [comment]. Science 280, 2074-2075.
- Hamilton, M.S., Ball, J., Bromidge, E., and Franklin, I.M. (1991).** Surface antigen expression of human neoplastic plasma cells includes molecules associated with lymphocyte recirculation and adhesion. Br. J. Haematol. 78, 60-65.
- Hahn, W. C., Y. Rosenstein, V. Calvo, S. J. Burakoff, and B. E. Bierer. (1992).** A distinct cytoplasmic domain of CD2 regulates ligand avidity and T-cell responsiveness to antigen. Proc. Natl. Acad. Sci. U. S. A 89:7179-7183.
- Han, J., Das, B., Wei, W., Van Aelst, L., Mosteller, R.D., Khosravi Far, R., Westwick, J.K., Der, C.J., and Broek, D. (1997).** Lck regulates Vav activation of members of the Rho family of GTPases. Mol. Cell Biol. 17, 1346-1353.
- Han, J., K. Luby Phelps, B. Das, X. Shu, Y. Xia, R. D. Mosteller, U. M. Krishna, J. R. Falck, M. A. White, and D. Broek. (1998).** Role of substrates and products of PI 3-kinase in regulating activation of Rac-related guanosine triphosphatases by Vav. Science 279:558-560.
- Hanke, J.H., Gardner, J.P., Dow, R.L., Changelian, P.S., Brissette, W.H., Weringer, E.J., Pollok, B.A., and Connelly, P.A. (1996).** Discovery of a novel, potent, and Src family-selective tyrosine kinase inhibitor. Study of Lck- and FynT-dependent T cell activation. J. Biol. Chem. 271, 695-701.
- Harder, T. and Simons, K. (1997).** Caveolae, DIGs, and the dynamics of sphingolipid-cholesterol microdomains. Curr. Opin. Cell Biol. 9, 534-542.
- Harder, T. and Simons, K. (1999).** Clusters of glycolipid and glycosylphosphatidylinositol-anchored proteins in lymphoid cells: accumulation of actin regulated by local tyrosine phosphorylation. Eur. J. Immunol. 29, 556-562.
- Harding, F.A., McArthur, J.G., Gross, J.A., Raulet, D.H., and Allison, J.P. (1992).** CD28-mediated signalling co-stimulates murine T cells and prevents induction of anergy in T-cell clones. Nature 356, 607-609.
- Hartwig, J. H., G. M. Bokoch, C. L. Carpenter, P. A. Janmey, L. A. Taylor, A. Toker, and T. P. Stossel. (1995).** Thrombin receptor ligation and activated Rac uncap actin filament barbed ends through phosphoinositide synthesis in permeabilized human platelets. Cell 82:643-653.
- Haynes, B.F., Telen, M.J., Hale, L.P., and Denning, S.M. (1989).** CD44--a molecule involved in leukocyte adherence and T-cell activation [published erratum appears in Immunol Today 1990 Mar;11(3):80]. Immunol. Today. 10, 423-428.
- He, Q., Lesley, J., Hyman, R., Ishihara, K., and Kincade, P.W. (1992).** Molecular isoforms of murine CD44 and evidence that the membrane proximal domain is not critical for hyaluronate recognition. J. Cell Biol. 119, 1711-1719.

- Herrlich, P., Zoller, M., Pals, S.T., and Ponta, H. (1993).** CD44 splice variants: metastases meet lymphocytes. *Immunol. Today.* 14, 395-399.
- Hirao, M., Sato, N., Kondo, T., Yonemura, S., Monden, M., Sasaki, T., Takai, Y., and Tsukita, S. (1996).** Regulation mechanism of ERM (ezrin/radixin/moesin) protein/plasma membrane association: possible involvement of phosphatidylinositol turnover and Rho-dependent signaling pathway. *J. Cell Biol.* 135, 37-51.
- Holdorf, A.D., Green, J.M., Levin, S.D., Denny, M.F., Straus, D.B., Link, V., Changelian, P.S., Allen, P.M., and Shaw, A.S. (1999).** Proline residues in CD28 and the Src homology (SH)3 domain of Lck are required for T cell costimulation. *J. Exp. Med.* 190, 375-384.
- Holsinger, L. J., I. A. Graef, W. Swat, T. Chi, D. M. Bautista, L. Davidson, R. S. Lewis, F. W. Alt, and G. R. Crabtree. (1998).** Defects in actin-cap formation in Vav-deficient mice implicate an actin requirement for lymphocyte signal transduction. *Curr. Biol.* 8:563-572.
- Huet, S., Groux, H., Caillou, B., Valentin, H., Prieur, A.M., and Bernard, A. (1989).** CD44 contributes to T cell activation. *J. Immunol.* 143, 798-801.
- Ilangumaran, S., Briol, A., and Hoessli, D.C. (1998).** CD44 selectively associates with active Src family protein tyrosine kinases Lck and Fyn in glycosphingolipid-rich plasma membrane domains of human peripheral blood lymphocytes. *Blood.* 91, 3901-3908.
- Ishiwatari Hayasaka, H., Fujimoto, T., Osawa, T., Hirama, T., Toyama Sorimachi, N., and Miyasaka, M. (1999).** Requirements for signal delivery through CD44: analysis using CD44-Fas chimeric proteins. *J. Immunol.* 163, 1258-1264.
- Izquierdo Pastor, M., Reif, K., and Cantrell, D. (1995).** The regulation and function of p21ras during T-cell activation and growth. *Immunol. Today.* 16, 159-164.
- Jackman, J.K., Motto, D.G., Sun, Q., Tanemoto, M., Turck, C.W., Peltz, G.A., Koretzky, G.A., and Findell, P.R. (1995).** Molecular cloning of SLP-76, a 76-kDa tyrosine phosphoprotein associated with Grb2 in T cells. *J. Biol. Chem.* 270, 7029-7032.
- Jain, J., Loh, C., and Rao, A. (1995).** Transcriptional regulation of the IL-2 gene. *Curr. Opin. Immunol.* 7, 333-342.
- Jalkanen, S., Jalkanen, M., Bargatze, R., Tammi, M., and Butcher, E.C. (1988).** Biochemical properties of glycoproteins involved in lymphocyte recognition of high endothelial venules in man. *J. Immunol.* 141, 1615-1623.
- Jalkanen, S. and Jalkanen, M. (1992).** Lymphocyte CD44 binds the COOH-terminal heparin-binding domain of fibronectin. *J. Cell Biol.* 116, 817-825.

- Janmey, P.A. (1998).** The cytoskeleton and cell signaling: component localization and mechanical coupling. *Physiol. Rev.* 78, 763-781.
- Janmey, P. A. and T. P. Stossel. (1989).** Gelsolin-polyphosphoinositide interaction. Full expression of gelsolin-inhibiting function by polyphosphoinositides in vesicular form and inactivation by dilution, aggregation, or masking of the inositol head group. *J Biol. Chem.* 264:4825-4831.
- Jenkins, M. K. (1994).** The ups and downs of T cell costimulation. *Immunity.* 1:443-446.
- Jenkinson, E.J., Kingston, R., Smith, C.A., Williams, G.T., and Owen, J.J. (1989).** Antigen-induced apoptosis in developing T cells: a mechanism for negative selection of the T cell receptor repertoire. *Eur. J. Immunol.* 19, 2175-2177.
- Jones, P.H., Bishop, L.A., and Watt, F.M. (1996).** Functional significance of CD9 association with beta 1 integrins in human epidermal keratinocytes. *Cell Adhes. Commun.* 4, 297-305.
- Ju, S-T., D. J. Panka, H. Cui, R. Ettinger, M. El-Khatib, D. H. Sherr, B. Z. Stanger, and A. Marshak-Rothstein. (1995).** Fas (CD95)/FasL interactions required for programmed cell death after T cell activation. *Nature* 373:444-448.
- Kaga, S., S. Ragg, K. A. Rogers, and A. Ochi. (1998).** Stimulation of CD28 with B7-2 promotes focal adhesion-like cell contacts where Rho family small G proteins accumulate in T cells. *J Immunol* 160:24-27.
- Kalomiris, E.L. and Bourguignon, L.Y. (1988).** Mouse T lymphoma cells contain a transmembrane glycoprotein (GP85) that binds ankyrin. *J. Cell Biol.* 106, 319-327.
- Kandzari, D.E., Chen, J., and Goldschmidt Clermont, P.J. (1996).** Regulation of the actin cytoskeleton by inositol phospholipid pathways. *Subcell. Biochem.* 26, 97-114.
- Kato, K., M. Koyanagi, H. Okada, T. Takanashi, Y. W. Wong, A. F. Williams, K. Okumura, and H. Yagita. (1992).** CD48 is a counter-receptor for mouse CD2 and is involved in T cell activation. *J Exp. Med.* 176:1241-1249.
- Katoh, S., Zheng, Z., Oritani, K., Shimozato, T., and Kincade, P.W. (1995).** Glycosylation of CD44 negatively regulates its recognition of hyaluronan. *J. Exp. Med.* 182, 419-429.
- Kersh, E.N., Shaw, A.S., and Allen, P.M. (1998).** Fidelity of T cell activation through multistep T cell receptor zeta phosphorylation [see comments]. *Science* 281, 572-575.
- Khaldoyanidi, S., Denzel, A., and Zoller, M. (1996).** Requirement for CD44 in proliferation and homing of hematopoietic precursor cells. *J. Leukoc. Biol.* 60, 579-592.

- Kobayashi, M., Imamura, M., Uede, T., Sakurada, K., Maeda, S., Iwasaki, H., Tsuda, Y., Musashi, M., and Miyazaki, T. (1994).** Expression of adhesion molecules on human hematopoietic progenitor cells at different maturational stages. *Stem. Cells. Dayt.* 12, 316-321.
- Kong Y., K. Fisher, M. F. Bachmann, S. Mariathasan, I. Kozieradzki, M. P. Nghiem, D. Bouchard, A. Bernstein, P. S. Ohashi, and J. M. Penniger. (1998).** Vav regulates peptide-specific apoptosis in thymocytes. *J. Exp. Med.* 188(11): 2099-2111
- Koopman, G., van Kooyk, Y., de Graaff, M., Meyer, C.J., Figdor, C.G., and Pals, S.T. (1990).** Triggering of the CD44 antigen on T lymphocytes promotes T cell adhesion through the LFA-1 pathway. *J. Immunol.* 145, 3589-3593.
- Kosugi, A., Saitoh, S.I., Noda, S., Yasuda, K., Hayashi, F., Ogata, M., and Hamaoka, T. (1999).** Translocation of tyrosine-phosphorylated TCR $\zeta$  chain to glycolipid-enriched membrane domains upon T cell activation. *Int. Immunol.* 11, 1395-1401.
- Kozma, R., Ahmed, S., Best, A., and Lim, L. (1995).** The Ras-related protein Cdc42Hs and bradykinin promote formation of peripheral actin microspikes and filopodia in Swiss 3T3 fibroblasts. *Mol. Cell Biol.* 15, 1942-1952.
- L mmli, U. K. (1970).** Cleavage of structural proteins during the assembly of the head of bacteriophage T4. *Nature* 227:680-685.
- Leeuwen, J. and Samelson, L.E. (1999).** T cell antigen-receptor signal transduction. *Curr. Opin. Immunol.* 11, 242-248.
- Lenardo, M.J., Boehme, S., Chen, L., Combadiere, B., Fisher, G., Freedman, M., McFarland, H., Pelfrey, C., and Zheng, L. (1995).** Autocrine feedback death and the regulation of mature T lymphocyte antigen responses. *Int. Rev. Immunol.* 13, 115-134.
- Lesley, J. and Hyman, R. (1992).** CD44 can be activated to function as an hyaluronic acid receptor in normal murine T cells. *Eur. J. Immunol.* 22, 2719-2723.
- Lesley, J., Kincade, P.W., and Hyman, R. (1993).** Antibody-induced activation of the hyaluronan receptor function of CD44 requires multivalent binding by antibody. *Eur. J. Immunol.* 23, 1902-1909.
- Liao, H.X., Levesque, M.C., Patton, K., Bergamo, B., Jones, D., Moody, M.A., Telen, M.J., and Haynes, B.F. (1993).** Regulation of human CD44H and CD44E isoform binding to hyaluronan by phorbol myristate acetate and anti-CD44 monoclonal and polyclonal antibodies. *J. Immunol.* 151, 6490-6499.
- Lin, J., Weiss, A., and Finco, T.S. (1999).** Localization of LAT in glycolipid-enriched microdomains is required for T cell activation. *J. Biol. Chem.* 274, 28861-28864.

- Lisanti, M.P., Scherer, P.E., Vidugiriene, J., Tang, Z., Hermanowski Vosatka, A., Tu, Y.H., Cook, R.F., and Sargiacomo, M. (1994).** Characterization of caveolin-rich membrane domains isolated from an endothelial-rich source: implications for human disease. *J. Cell Biol.* 126, 111-126.
- Liu, D., Zhang, D., Mori, H., and Sy, M.S. (1996).** Binding of CD44 to hyaluronic acid can be induced by multiple signals and requires the CD44 cytoplasmic domain. *Cell Immunol.* 174, 73-83.
- Lokeshwar, V.B. and Bourguignon, L.Y. (1991).** Post-translational protein modification and expression of ankyrin-binding site(s) in GP85 (Pgp-1/CD44) and its biosynthetic precursors during T-lymphoma membrane biosynthesis. *J. Biol. Chem.* 266, 17983-17989.
- Lokeshwar, V.B. and Bourguignon, L.Y. (1992).** Tyrosine phosphatase activity of lymphoma CD45 (GP180) is regulated by a direct interaction with the cytoskeleton. *J. Biol. Chem.* 267:21551-21557.
- Lokeshwar, V.B., Fregien, N., and Bourguignon, L.Y. (1994).** Ankyrin-binding domain of CD44(GP85) is required for the expression of hyaluronic acid-mediated adhesion function. *J. Cell Biol.* 126, 1099-1109.
- Lokeshwar, V.B., Iida, N., and Bourguignon, L.Y. (1996).** The cell adhesion molecule, GP116, is a new CD44 variant (ex14/v10) involved in hyaluronic acid binding and endothelial cell proliferation. *J. Biol. Chem.* 271, 23853-23864.
- Lowin Kropf, B., V. S. Shapiro, and A. Weiss. (1998).** Cytoskeletal polarization of T cells is regulated by an immunoreceptor tyrosine-based activation motif-dependent mechanism. *J. Cell Biol.* 140:861-871.
- Mackay, D.J., Esch, F., Furthmayr, H., and Hall, A. (1997).** Rho- and rac-dependent assembly of focal adhesion complexes and actin filaments in permeabilized fibroblasts: an essential role for ezrin/radixin/moesin proteins. *J. Cell Biol.* 138, 927-938.
- Mackay, D.J. and Hall, A. (1998).** Rho GTPases. *J. Biol. Chem.* 273, 20685-20688.
- Maecker, H.T., Todd, S.C., and Levy, S. (1997).** The tetraspanin superfamily: molecular facilitators. *FASEB J.* 11, 428-442.
- Maness, P. F. and R. C. Walsh. (1982).** Dihydrocytochalasin B disorganizes actin cytoarchitecture and inhibits initiation of DNA synthesis in 3T3 cells. *Cell* 30:253-262.
- Manser, E., Leung, T., Salihuddin, H., Zhao, Z.S., and Lim, L. (1994).** A brain serine/threonine protein kinase activated by Cdc42 and Rac1. *Nature* 367, 40-46.
- Matsumoto, G., Nghiem, M.P., Nozaki, N., Schmits, R., and Penninger, J.M. (1998).** Cooperation between CD44 and LFA-1/CD11a adhesion receptors in lymphokine-activated killer cell cytotoxicity. *J. Immunol.* 160, 5781-5789.

- Matsuyama, T., A. Yamada, K. Deusch, J. Sleasman, J. F. Daley, Y. Torimoto, and T. Abe. (1991).** Cytochalasins enhance the proliferation of CD4 cells through the CD3-Ti antigen receptor complex or the CD2 molecule through an effect on early events of activation. *J Immunol* 146:3736-3741.
- McConkey, D.J., Fosdick, L., D'Adamio, L., Jondal, M., and Orrenius, S. (1994).** Co-receptor (CD4/CD8) engagement enhances CD3-induced apoptosis in thymocytes. Implications for negative selection. *J. Immunol.* 153, 2436-2443.
- Meng, F. and Lowell, C.A. (1998).** A beta 1 integrin signaling pathway involving Src-family kinases, Cbl and PI-3 kinase is required for macrophage spreading and migration. *EMBO J.* 17, 4391-4403.
- Michel, F., Grimaud, L., Tuosto, L., and Acuto, O. (1998).** Fyn and ZAP-70 are required for Vav phosphorylation in T cells stimulated by antigen-presenting cells. *J. Biol. Chem.* 273, 31932-31938.
- Miki, H., K. Miura, and T. Takenawa. (1996).** N-WASP, a novel actin-depolymerizing protein, regulates the cortical cytoskeletal rearrangement in a PIP2-dependent manner downstream of tyrosine kinases. *EMBO J* 15:5326-5335.
- Miyake, K., Medina, K.L., Hayashi, S., Ono, S., Hamaoka, T., and Kincade, P.W. (1990).** Monoclonal antibodies to Pgp-1/CD44 block lympho-hemopoiesis in long-term bone marrow cultures. *J. Exp. Med.* 171, 477-488.
- Montixi, C., Langlet, C., Bernard, A.M., Thimonier, J., Dubois, C., Wurbel, M.A., Chauvin, J.P., Pierres, M., and He, H.T. (1998).** Engagement of T cell receptor triggers its recruitment to low-density detergent-insoluble membrane domains. *EMBO J.* 17, 5334-5348.
- Moran, M. and Miceli, M. C. (1998).** Engagement of GPI-linked CD48 contributes to TCR signals and cytoskeletal reorganization: A role for lipid rafts in T cell activation. *Immunity* 9:787-796
- Morimoto, K., Robin, E., Le Bousse Kerdiles, M.C., Li, Y., Clay, D., Jasmin, C., and Smadja Joffe, F. (1994).** CD44 mediates hyaluronan binding by human myeloid KG1A and KG1 cells. *Blood.* 83, 657-662.
- Naor, D., Sionov, R.V., Zahalka, M., Rochman, M., Holzmann, B., and Ish Shalom, D. (1998).** Organ-specific requirements for cell adhesion molecules during lymphoma cell dissemination. *Curr. Top. Microbiol. Immunol.* 231, 143-166.
- Naujokas, M.F., Morin, M., Anderson, M.S., Peterson, M., and Miller, J. (1993).** The chondroitin sulfate form of invariant chain can enhance stimulation of T cell responses through interaction with CD44. *Cell* 74, 257-268.

- Neame, S.J. and Isacke, C.M. (1992).** Phosphorylation of CD44 in vivo requires both Ser323 and Ser325, but does not regulate membrane localization or cytoskeletal interaction in epithelial cells. *EMBO J.* 11, 4733-4738.
- Neame, S.J. and Isacke, C.M. (1993).** The cytoplasmic tail of CD44 is required for basolateral localization in epithelial MDCK cells but does not mediate association with the detergent-insoluble cytoskeleton of fibroblasts. *J. Cell Biol.* 121, 1299-1310.
- Nel, A.E., Gupta, S., Lee, L., Ledbetter, J.A., and Kanner, S.B. (1995).** Ligation of the T-cell antigen receptor (TCR) induces association of hSos1, ZAP-70, phospholipase C-gamma 1, and other phosphoproteins with Grb2 and the zeta-chain of the TCR. *J. Biol. Chem.* 270, 18428-18436.
- Nicoletti, I., Migliorati, G., Pagliacci, M.C., Grignani, F., and Riccardi, C. (1991).** A rapid and simple method for measuring thymocyte apoptosis by propidium iodide staining and flow cytometry. *J. Immunol. Methods* 139, 271-279.
- Nobes, C.D. and Hall, A. (1995).** Rho, Rac and Cdc42 GTPases Regulate the Assembly of Multimolecular Focal Complexes Associated with Actin Stress Fibers, Lamellipodia, and Filopodia. *Cell* 81, 53-62.
- Nunes, J.A., Truneh, A., Olive, D., and Cantrell, D.A. (1996).** Signal transduction by CD28 costimulatory receptor on T cells. B7-1 and B7-2 regulation of tyrosine kinase adaptor molecules. *J. Biol. Chem.* 271, 1591-1598.
- O'Rourke, A.M., Apgar, J.R., Kane, K.P., Martz, E., and Mescher, M.F. (1991).** Cytoskeletal function in CD8- and T cell receptor-mediated interaction of cytotoxic T lymphocytes with class I protein. *J. Exp. Med.* 173, 241-249.
- Oliferenko, S., Paiha, K., Harder, T., Gerke, V., Schwarzler, C., Schwarz, H., Beug, H., Gunthert, U., and Huber, L.A. (1999).** Analysis of CD44-containing lipid rafts: Recruitment of annexin II and stabilization by the actin cytoskeleton. *J. Cell Biol.* 146, 843-854.
- Olson, M.F., Pasteris, N.G., Gorski, J.L., and Hall, A. (1996).** Faciogenital dysplasia protein (FGD1) and Vav, two related proteins required for normal embryonic development, are upstream regulators of Rho GTPases. *Curr. Biol.* 6, 1628-1633.
- Pardi, R., L. Inverardi, C. Rugarli, and J. R. Bender. (1992).** Antigen-receptor complex stimulation triggers protein kinase C-dependent CD11a/CD18-cytoskeleton association in T lymphocytes. *J Cell Biol.* 116:1211-1220.
- Peach, R.J., Hollenbaugh, D., Stamenkovic, I., and Aruffo, A. (1993).** Identification of hyaluronic acid binding sites in the extracellular domain of CD44. *J. Cell Biol.* 122, 257-264.
- Penninger, J.M. and Crabtree, G.R. (1999).** The actin cytoskeleton and lymphocyte activation. *Cell* 96, 9-12.

- Perlmutter, R. M., S. D. Levin, M. W. Appleby, S. J. Anderson, and J. Alberola-Ila. (1993).** Regulation of lymphocyte function by protein phosphorylation. Annual reviews in immunology 11:451-499.
- Price, L.S., Leng, J., Schwartz, M.A., and Bokoch, G.M. (1998).** Activation of Rac and Cdc42 by integrins mediates cell spreading. Mol. Biol. Cell 9, 1863-1871.
- Punt, J.A., Havran, W., Abe, R., Sarin, A., and Singer, A. (1997).** T cell receptor (TCR)-induced death of immature CD4+CD8+ thymocytes by two distinct mechanisms differing in their requirement for CD28 costimulation: implications for negative selection in the thymus. J. Exp. Med. 186, 1911-1922.
- Qian, D. and Weiss, A. (1997).** T cell antigen receptor signal transduction. Curr. Opin. Cell Biol. 9, 205-212.
- Raab, M., da Silva, A.J., Findell, P.R., and Rudd, C.E. (1997).** Regulation of Vav-SLP-76 binding by ZAP-70 and its relevance to TCR zeta/CD3 induction of interleukin-2. Immunity. 6, 155-164.
- Rafi, A., Nagarkatti, M., and Nagarkatti, P.S. (1997).** Hyaluronate-CD44 interactions can induce murine B-cell activation. Blood. 89, 2901-2908.
- Rajnavolgyi, E., Nagy, Z., Kurucz, I., Gogolak, P., Toth, G.K., Varadi, G., Penke, B., Tigy, Z., Hollosi, M., and Gergely, J. (1994).** T cell recognition of the posttranslationally cleaved intersubunit region of influenza virus hemagglutinin. Mol. Immunol. 31, 1403-1414.
- Ridley, A.J. and Hall, A. (1992).** Distinct patterns of actin organization regulated by the small GTP-binding proteins Rac and Rho. Cold. Spring. Harb. Symp. Quant. Biol. 57, 661-671.
- Ridley, A.J., Paterson, H.F., Johnston, C.L., Diekmann, D., and Hall, A. (1992).** The small GTP-binding protein rac regulates growth factor-induced membrane ruffling. Cell 70, 401-410.
- Rincon, M. and Flavell, R.A. (1994).** AP-1 transcriptional activity requires both T-cell receptor-mediated and co-stimulatory signals in primary T lymphocytes. EMBO J. 13, 4370-4381.
- Rozdzial, M. M., B. Malissen, and T. H. Finkel. (1995).** Tyrosine-phosphorylated T cell receptor zeta chain associates with the actin cytoskeleton upon activation of mature T lymphocytes. Immunity. 3:623-633.
- Rosel, M., Foger, N., and Zoller, M. (1998).** Involvement of CD44 exon v10 in B-cell activation. Tissue. Antigens. 52, 99-113.

- Rothenberg, E.V. and Ward, S.B. (1996).** A dynamic assembly of diverse transcription factors integrates activation and cell-type information for interleukin 2 gene regulation. Proc. Natl. Acad. Sci. U. S. A. 93, 9358-9365.
- Rothman, B.L., Blue, M.L., Kelley, K.A., Wunderlich, D., Mierz, D.V., and Aune, T.M. (1991).** Human T cell activation by OKT3 is inhibited by a monoclonal antibody to CD44. J. Immunol. 147, 2493-2499.
- Rozsnyay, Z. (1999).** Signaling complex formation of CD44 with src-related kinases. Immunol. Lett. 68, 101-108.
- Ryser, J. E., E. Rungger Brandle, C. Chaponnier, G. Gabbiani, and P. Vassalli. (1982).** The area of attachment of cytotoxic T lymphocytes to their target cells shows high motility and polarization of actin, but not myosin. J Immunol 128:1159-1162.
- Sancho, J., Peter, M.E., Franco, R., Danielian, S., Kang, J.S., Fagard, R., Woods, J., Reed, J.C., Kamoun, M., and Terhorst, C. (1993).** Coupling of GTP-binding to the T cell receptor (TCR) zeta-chain with TCR-mediated signal transduction. J. Immunol. 150, 3230-3242.
- Scheeren, R.A., Koopman, G., Van der Baan, S., Meijer, C.J., and Pals, S.T. (1991).** Adhesion receptors involved in clustering of blood dendritic cells and T lymphocytes. Eur. J. Immunol. 21, 1101-1105.
- Schroeder, R.J., Ahmed, S.N., Zhu, Y., London, E., and Brown, D.A. (1998).** Cholesterol and sphingolipid enhance the Triton X-100 insolubility of glycosylphosphatidylinositol-anchored proteins by promoting the formation of detergent-insoluble ordered membrane domains. J. Biol. Chem. 273, 1150-1157.
- Schwartz, L.M. (1992).** Insect muscle as a model for programmed cell death. J. Neurobiol. 23, 1312-1326.
- Sconocchia, G., Titus, J.A., and Segal, D.M. (1994).** CD44 is a cytotoxic triggering molecule in human peripheral blood NK cells. J. Immunol. 153, 5473-5481.
- Screaton, G.R., Bell, M.V., Jackson, D.G., Cornelis, F.B., Gerth, U., and Bell, J.I. (1992).** Genomic structure of DNA encoding the lymphocyte homing receptor CD44 reveals at least 12 alternatively spliced exons. Proc. Natl. Acad. Sci. U. S. A. 89, 12160-12164.
- Sechrist, J.P., Karnitz, L., and Abraham, R.T. (1991).** T-cell antigen receptor ligation induces tyrosine phosphorylation of phospholipase C-g1. J. Biol. Chem. 266, 12135-12139.
- Seth, A., Gote, L., Nagarkatti, M., Nagarkatti, P.S. (1991).** T cell receptor independent activation of cytolytic activity of cytotoxic T lymphocytes mediated through CD44 and gp90MEL-14. Proc. Natl. Acad. Sci. USA 88, 7877-7881.

- Shahinian, A., Pfeffer, K., Lee, K.P., Kundig, T.M., Kishihara, K., Wakeham, A., Kawai, K., Ohashi, P.S., Thompson, C.B., and Mak, T.W. (1993).** Differential T cell costimulatory requirements in CD28-deficient mice. *Science* 261, 609-612.
- Shaw, A.S. and Dustin, M.L.. (1997).** Making the T cell receptor go the distance: a topological view of T cell activation. *Immunity*. 6:361-369.
- Shaw, S. and Shimizu, Y. (1988).** Two molecular pathways of human T cell adhesion: establishment of receptor-ligand relationship. *Curr. Opin. Immunol* 1:92-97.
- Shimizu, Y., van Seventer, G.A., Siraganian, R., Wahl, L., and Shaw, S. (1989).** Dual role of the CD44 molecule in T cell adhesion and activation. *J. Immunol.* 143, 2457-2463.
- Shimizu, Y., J. L. Mobley, L. D. Finkelstein, and A. S. Chan. (1995).** A role for phosphatidylinositol 3-kinase in the regulation of beta 1 integrin activity by the CD2 antigen. *J Cell Biol.* 131:1867-1880.
- Shimizu, Y. and Shaw, S. (1991).** Lymphocyte interactions with extracellular matrix. *FASEB J.* 5, 2292-2299.
- Shimizu, Y., van Seventer, G.A., Ennis, E., Newman, W., Horgan, K.J., and Shaw, S. (1992).** Crosslinking of the T cell-specific accessory molecules CD7 and CD28 modulates T cell adhesion. *J. Exp. Med.* 175, 577-582.
- Simons, K. and Ikonen, E. (1997).** Functional rafts in cell membranes. *Nature* 387, 569-572.
- Sloan Lancaster, J., Shaw, A.S., Rothbard, J.B., and Allen, P.M. (1994).** Partial T cell signaling: altered phospho-zeta and lack of zap70 recruitment in APL-induced T cell anergy [see comments]. *Cell* 79, 913-922.
- Smith, C.A., Williams, G.T., Kingston, R., Jenkinson, E.J., and Owen, J.J.T. (1989).** Antibodies to CD3/T-cell receptor complex induce death by apoptosis in immature T cells in thymic cultures. *Nature* 337, 181-184.
- Snapper, S. B., F. S. Rosen, E. Mizoguchi, P. Cohen, W. Khan, C. H. Liu, T. L. Hagemann, S. P. Kwan, R. Ferrini, L. Davidson, A. K. Bhan, and F. W. Alt. (1998).** Wiskott-Aldrich syndrome protein-deficient mice reveal a role for WASP in T but not B cell activation. *Immunity*. 9:81-91.
- Sommer, F., Huber, M., Rollinghoff, M., and Lohoff, M. (1995).** CD44 plays a co-stimulatory role in murine T cell activation: ligation of CD44 selectively co-stimulates IL-2 production, but not proliferation in TCR-stimulated murine Th1 cells. *Int. Immunol.* 7, 1779-1786.
- Soula, M., Rothhut, B., Camoin, L., Guillaume, J.L., Strosberg, D., Vorherr, T., Burn, P., Meggio, F., Fischer, S., and Fagard, R. (1993).** Anti-CD3 and phorbol ester induce distinct phosphorylated sites in the SH2 domain of p56lck. *J. Biol. Chem.* 268, 27420-27427.

- Sperling, A.I. and Bluestone, J.A. (1996).** The complexities of T-cell co-stimulation: CD28 and beyond. *Immunol. Rev.* 153, 155-182.
- Springer, T.A., Dustin, M.L., Kishimoto, T.K., and Marlin, S.D. (1987).** The lymphocyte function-associated LFA-1, CD2, and LFA-3 molecules: cell adhesion receptors of the immune system. *Annu. Rev. Immunol.* 5, 223-252.
- Springer, T.A. (1994).** Traffic signals for lymphocyte recirculation and leukocyte emigration: the multistep paradigm. *Cell* 76, 301-314.
- St.John, T., Meyer, J., Idzerda, R., and Gallatin, W.M. (1990).** Expression of CD44 confers a new adhesive phenotype on transfected cells. *Cell* 60, 45-52.
- Stamenkovic, I., Aruffo, A., Amiot, M., and Seed, B. (1991).** The hematopoietic and epithelial forms of CD44 are distinct polypeptides with different adhesion potentials for hyaluronate-bearing cells. *EMBO J.* 10, 343-348.
- Stevens, R.L., Avraham, S., Gartner, M.C., Bruns, G.A., Austen, K.F., and Weis, J.H. (1988).** Isolation and characterization of a cDNA that encodes the peptide core of the secretory granule proteoglycan of human promyelocytic leukemia HL-60 cells. *J. Biol. Chem.* 263, 7287-7291.
- Stossel, T. P. (1994).** The machinery of cell crawling. *Sci. Am.* 271:54-5, 58.
- Straus, D.B. and Weiss, A. (1992).** Genetic evidence for the involvement of the lck tyrosine kinase in signal transduction through the T cell antigen receptor. *Cell* 70, 585-593.
- Su, B., Jacinto, E., Hibi, M., Kallunki, T., Karin, M., and Ben Neriah, Y. (1994).** JNK is involved in signal integration during costimulation of T lymphocytes. *Cell* 77, 727-736.
- Sy, M.S., Liu, D., Schiavone, R., Ma, J., Mori, H., and Guo, Y. (1996).** Interactions between CD44 and hyaluronic acid: their role in tumor growth and metastasis. *Curr. Top. Microbiol. Immunol.* 213, 129-153.
- Taher, T.E., Smit, L., Griffioen, A.W., Schilder Tol, E.J., Borst, J., and Pals, S.T. (1996).** Signaling through CD44 is mediated by tyrosine kinases. Association with p56lck in T lymphocytes. *J. Biol. Chem.* 271, 2863-2867.
- Tapon, N., Nagata, K., Lamarche, N., and Hall, A. (1998).** A new rac target POSH is an SH3-containing scaffold protein involved in the JNK and NF-kappaB signalling pathways. *EMBO J.* 17, 1395-1404.
- Teramoto, H., Crespo, P., Coso, O.A., Igishi, T., Xu, N., and Gutkind, J.S. (1996).** The small GTP-binding protein rho activates c-Jun N-terminal kinases/stress-activated protein kinases in human kidney 293T cells. Evidence for a Pak-independent signaling pathway. *J. Biol. Chem.* 271, 25731-25734.

- Thomas, M.L. (1999).** The regulation of antigen-receptor signaling by protein tyrosine phosphatases: a hole in the story. *Curr. Opin. Immunol.* 11, 270-276.
- Thomas, S.M., Soriano, P., and Imamoto, A. (1995).** Specific and redundant roles of Src and Fyn in organizing the cytoskeleton. *Nature* 376, 267-271.
- Thompson, C. B. 1995.** Distinct roles for the costimulatory ligands B7-1 and B7-2 in T helper cell differentiation? *Cell* 81:979-982.
- Tolg, C., Hofmann, M., Herrlich, P., and Ponta, H. (1993).** Splicing choice from ten variant exons establishes CD44 variability. *Nucleic Acids Res.* 21, 1225-1229.
- Toyama Sorimachi, N. and Miyasaka, M. (1994).** A sulfated proteoglycan as a novel ligand for CD44. *J. Dermatol.* 21, 795-801.
- Toyama Sorimachi, N., Sorimachi, H., Tobita, Y., Kitamura, F., Yagita, H., Suzuki, K., and Miyasaka, M. (1995).** A novel ligand for CD44 is serglycin, a hematopoietic cell lineage-specific proteoglycan. Possible involvement in lymphoid cell adherence and activation. *J. Biol. Chem.* 270, 7437-7444.
- Traunecker, A., Dolder, B., and Karjalainen, K. (1986).** A novel approach for preparing anti-T cell receptor constant region antibodies. *Eur. J. Immunol.* 16, 851-854.
- Tsukita, S., Oishi, K., Sato, N., Sagara, J., and Kawai, A. (1994).** ERM family members as molecular linkers between the cell surface glycoprotein CD44 and actin-based cytoskeletons. *J. Cell Biol.* 126, 391-401.
- Tsukita, S. and Yonemura, S. (1997).** ERM proteins: head-to-tail regulation of actin-plasma membrane interaction. *Trends Biochem. Sci.* 22, 53-58.
- Tuosto, L., and Acuto, O. (1998).** CD28 affects the earliest signaling events generated by TCR engagement. *Eur. J Immunol.* 28:2131-2142.
- Turley, E.A., Austen, L., Vandeligt, K., and Clary, C. (1991).** Hyaluronan and a cell-associated hyaluronan binding protein regulate the locomotion of ras-transformed cells. *J. Cell Biol.* 112, 1041-1047.
- Valitutti, S., M. Dessing, K. Aktories, H. Gallati, and A. Lanzavecchia. (1995).** Sustained signaling leading to T cell activation results from prolonged T cell receptor occupancy. Role of T cell actin cytoskeleton. *J Exp. Med.* 181:577-584.
- van Leeuwen, J.E. and Samelson, L.E. (1999).** T cell antigen-receptor signal transduction. *Curr. Opin. Immunol.* 11, 242-248.

- van Oers, N.S., Lowin Kropf, B., Finlay, D., Connolly, K., and Weiss, A. (1996).** alpha beta T cell development is abolished in mice lacking both Lck and Fyn protein tyrosine kinases. *Immunity.* 5, 429-436.
- Varma, R. and Mayor, S. (1998).** GPI-anchored proteins are organized in submicron domains at the cell surface. *Nature* 394, 798-801.
- Veillette, A., Zuniga Pflucker, J.C., Bolen, J.B., and Kruisbeek, A.M. (1989).** Engagement of CD4 and CD8 expressed on immature thymocytes induces activation of intracellular tyrosine phosphorylation pathways. *J. Exp. Med.* 170, 1671-1680.
- Verfaillie, C.M., Benis, A., Iida, J., McGlave, P.B., and McCarthy, J.B. (1994).** Adhesion of committed human hematopoietic progenitors to synthetic peptides from the C-terminal heparin-binding domain of fibronectin: cooperation between the integrin alpha 4 beta 1 and the CD44 adhesion receptor. *Blood.* 84, 1802-1811.
- Vermot Desroches, C., Wijdenes, J., Valmu, L., Roy, C., Pigott, R., Nortamo, P., and Gahmberg, C.G. (1995).** A CD44 monoclonal antibody differentially regulates CD11a/CD18 binding to intercellular adhesion molecules CD54, CD102 and CD50. *Eur. J. Immunol.* 25, 2460-2464.
- Viola, A., Schroeder, S., Sakakibara, Y. and A. Lanzavecchia. 1999.** T lymphocyte costimulation mediated by reorganization of membrane microdomains. *Science* 283, 680-682
- Wang, W., Gulden, P.H., Pierce, R.A., Shabanowitz, J.A., Man, S.T., Hunt, D.F., and Engelhard, V.H. (1997).** A naturally processed peptide presented by HLA-A\*0201 is expressed at low abundance and recognized by an alloreactive CD8+ cytotoxic T cell with apparent high affinity. *J. Immunol.* 158, 5797-5804.
- Wange, R.L. and Samelson, L.E.. (1996).** Complex complexes: Signaling at the TCR. *Immunity* 5:197-205.
- Waterman-Storer, C.M., Worthylake, R.A., Liu, B.P., Burridge, K., and Salmon, E.D. (1999).** Microtubule growth activates Rac1 to promote lamellipodial protrusion in fibroblasts. *Nature* 1, 45-50.
- Watts, J.D., Welham, M.J., Kalt, L., Schrader, J.W., and Aebersold, R. (1993).** IL-2 stimulation of T lymphocytes induces sequential activation of mitogen-activated protein kinases and phosphorylation of p56lck at serine-59. *J. Immunol.* 151, 6862-6871.
- Weber, G.F., Ashkar, S., Glimcher, M.J., and Cantor, H. (1996).** Receptor-ligand interaction between CD44 and osteopontin (Eta-1). *Science* 271, 509-512.
- Weiss, A. (1993).** T cell antigen receptor signal transduction: A tale of tails and cytoplasmic protein-tyrosine kinases. *Cell* 73:209-212.

- Weiss, A., Koretzky, G., Schatzman, R.C., and Kadlecak, T. (1991).** Functional activation of the T-cell antigen receptor induces tyrosine phosphorylation of phospholipase C-gamma 1. Proc. Natl. Acad. Sci. U. S. A. 88, 5484-5488.
- Weiss, A. and Littman, D.R. (1994).** Signal transduction by lymphocyte antigen receptors. Cell 76:263-274.
- Wiest, D.L., Ashe, J.M., Abe, R., Bolen, J.B., and Singer, A. (1996).** TCR activation of ZAP70 is impaired in CD4+CD8+ thymocytes as a consequence of intrathymic interactions that diminish available p56lck. Immunity. 4, 495-504.
- Winkler, D.G., Park, I., Kim, T., Payne, N.S., Walsh, C.T., Strominger, J.L., and Shin, J. (1993).** Phosphorylation of Ser-42 and Ser-59 in the N-terminal region of the tyrosine kinase p56lck. Proc. Natl. Acad. Sci. U. S. A. 90, 5176-5180.
- Winoto, A. (1997).** Cell death in the regulation of immune responses. Curr. Opin. Immunol. 9, 365-370.
- Woscholski, R., T. Kodaki, M. McKinnon, M. D. Waterfield, and P. J. Parker. (1994).** A comparison of demethoxyviridin and wortmannin as inhibitors of phosphatidylinositol 3-kinase. FEBS Lett. 342:109-114.
- W Ifling, C. and M. M. Davis. 1998.** A receptor/cytoskeletal movement triggered by costimulation during T cell activation. Science 282:2266-2269.
- Wu, L., Kincade, P.W., and Shortman, K. (1993).** The CD44 expressed on the earliest intrathymic precursor population functions as a thymus homing molecule but does not bind to hyaluronate. Immunol. Lett. 38, 69-75.
- Wyllie, A.H. (1980).** Glucocorticoid-induced thymocyte apoptosis is associated with endogenous endonuclease activation. Nature 284, 555-556.
- Xavier, R., Brennan, T., Li, Q., McCormack, C., and Seed, B. (1998).** Membrane compartmentation is required for efficient T cell activation. Immunity. 8, 723-732.
- Xu, H. and Littman, D.R. (1993).** A kinase-independent function of Lck in potentiating antigen-specific T cell activation. Cell 74, 633-643.
- Yablonski, D., Kuhne, M.R., Kadlecak, T., and Weiss, A. (1998).** Uncoupling of nonreceptor tyrosine kinases from PLC-gamma1 in an SLP-76-deficient T cell. Science 281, 413-416.
- Yang, Y., Mercep, M., Ware, C.F. and Ashwell, J.D. (1995).** Fas and activation-induced Fas ligand mediate apoptosis of T cell hybridomas: inhibition of Fas ligand expression by retinoic acid and glucocorticoids. J Exp Med. 181:1673-1682.

- Yashiro, Y., Tai, X.G., Toyo oka, K., Park, C.S., Abe, R., Hamaoka, T., Kobayashi, M., Neben, S., and Fujiwara, H. (1998).** A fundamental difference in the capacity to induce proliferation of naive T cells between CD28 and other co-stimulatory molecules. *Eur. J. Immunol.* 28, 926-935.
- Yu, Q. and Toole, B.P. (1996).** A new alternatively spliced exon between v9 and v10 provides a molecular basis for synthesis of soluble CD44. *J. Biol. Chem.* 271, 20603-20607.
- Zacharchuk, C.M., Mercep, M., Chakraborti, P.K., Simons, S.S., Jr., and Ashwell, J.D. (1990).** Programmed T lymphocyte death. Cell activation- and steroid-induced pathways are mutually antagonistic. *J. Immunol.* 145, 4037-4045.
- Zawadzki, V., Perschl, A., Rosel, M., Hekele, A., and Zoller, M. (1998).** Blockade of metastasis formation by CD44-receptor globulin. *Int. J. Cancer* 75, 919-924.
- Zhang, W., Sloan Lancaster, J., Kitchen, J., Trible, R.P., and Samelson, L.E. (1998a).** LAT: the ZAP-70 tyrosine kinase substrate that links T cell receptor to cellular activation. *Cell* 92, 83-92.
- Zhang, W., Trible, R.P., and Samelson, L.E. (1998b).** LAT palmitoylation: its essential role in membrane microdomain targeting and tyrosine phosphorylation during T cell activation. *Immunity*. 9, 239-246.
- Zheng, Z., Katoh, S., He, Q., Oritani, K., Miyake, K., Lesley, J., Hyman, R., Hamik, A., Parkhouse, R.M., Farr, A.G., and et al, (1995).** Monoclonal antibodies to CD44 and their influence on hyaluronan recognition. *J. Cell Biol.* 130, 485-495.
- Zhou, D.F., Ding, J.F., Picker, L.J., Bargatze, R.F., Butcher, E.C., and Goeddel, D.V. (1989).** Molecular cloning and expression of Pgp-1. The mouse homolog of the human H-CAM (Hermes) lymphocyte homing receptor. *J. Immunol.* 143, 3390-3395.
- Zoller, M. (1995).** CD44: physiological expression of distinct isoforms as evidence for organ-specific metastasis formation. *J. Mol. Med.* 73, 425-438.
- Zoller, M. (1996).** Joint features of metastasis formation and lymphocyte maturation and activation. *Curr. Top. Microbiol. Immunol.* 213, 215-247.
- Zuckerman, L.A., Pullen, L., and Miller, J. (1998).** Functional consequences of costimulation by ICAM-1 on IL-2 gene expression and T cell activation. *J. Immunol.* 160, 3259-3268.