

- Ma H.-T., Venkatachalam K., Parys J.B., Gill D.L.* 2002. Modification of store-operated channel coupling and inositol trisphosphate receptor function by 2-aminoethoxydiphenyl borate in DT40 lymphocytes. *J. Biol. Chem.* 277 : 6915–6922.
- McElroy S.P., Gurney A.M., Drummond R.M.* 2008. Pharmacological profile of store-operated Ca^{2+} entry in intrapulmonary artery smooth muscle cells. *Eur. J. Pharmacol.* 584 : 10–20.
- Monahan R.A., Dvorak H.F., Dvorak A.M.* 1981. Ultrastructural localization of nonspecific esterase activity in guinea pig and human monocytes, macrophages and lymphocytes. *Blood.* 58 : 1089–1099.
- Moreno C., Vaca L.* 2012. Microdomain organization of SOCE signaling. In: *Store-operated Ca^{2+} entry (SOCE) pathways*. Wien, Springer-Verlag. 93–113.
- Nguyen N.T., Han W., Cao W.-M., Wang Y., Wen S., Huang Y., Li M., Du L., Zhou Y.* 2018. Store-operated calcium entry mediated by Orai and STIM. *Comprehensive Physiol.* 8 : 981–1002.
- Nurbaeva M.K., Eckstein M., Snead M.L., Feske S., Lacruz R.S.* 2015. Store-operated Ca^{2+} entry modulates the expression of enamel genes. *J. Dent. Res.* 94 : 1471–1477.
- Nwokonko R.M., Cai X., Loktionova N.A., Wang Y., Zhou Y., Gill D.L.* 2017. The STIM–Orai pathway: Conformational coupling between STIM and Orai in the activation of store-operated Ca^{2+} entry. *Adv. Exp. Med. Biol.* 993 : 83–98.
- Ohana L., Newell E.W., Stanley E.F., Schlichter L.C.* 2009. The Ca^{2+} release-activated Ca^{2+} current (I_{CRAC}) mediates store-operated Ca^{2+} entry in rat microglia. *Channels.* 3 : 129–139.
- Peinelt C., Lis A., Beck A., Fleig A., Penner R.* 2008. 2-aminoethoxydiphenyl borate directly facilitates and indirectly inhibits STIM1-dependent gating of CRAC channels. *J. Physiol.* 586 : 3061–3073.
- Prakriya M., Lewis R.S.* 2001. Potentiation and inhibition of Ca^{2+} release-activated Ca^{2+} channels by 2-aminoethoxydiphenyl borate (2-APB) occurs independently of IP_3 receptors. *J. Physiol.* 536 : 3–19.
- Prakriya M., Lewis R.S.* 2015. Store-operated calcium channels. *Physiol. Rev.* 95 : 1383–1436.
- Putney J.W.* 1990. Capacitative calcium entry revisited. *Cell Calcium.* 11 : 611–624.
- Putney J.W.* 2001. Pharmacology of capacitative calcium entry. *Mol. Interv.* 1 : 84–94.
- Putney J.W.* 2010. Pharmacology of store-operated calcium channels. *Mol. Interv.* 10 : 209–218.
- Putney J.W.* 2011. The physiological function of store-operated calcium entry. *Neurochem. Res.* 36 : 1157–1165.
- Putney J.W.* 2017. Store-operated calcium entry: An historical overview. *Adv. Exp. Med. Biol.* 981 : 205–214.
- Randriamampita C., Trautmann A.* 1987. Ionic channels in murine macrophages. *Cell. Biol.* 105 : 761–769.
- Shaw P.J., Feske S.* 2013. Physiological and pathophysiological functions of SOCE in the immune system. *Front. Biosci.* 4 : 2253–2268.
- Thastrup O., Dawson A.P., Scharff O., Foder B., Cullen P.J., Drobak B.K., Bjerrum P.J., Christensen S.B., Hanley M.R.* 1989. Thapsigargin, a novel molecular probe for studying intracellular calcium release and storage. *Agents Actions.* 27 : 17–23.
- Vaca L.* 2010. SOCIC: The store-operated calcium influx complex. *Cell Calcium.* 47 : 199–209.
- Vaeth M., Zee I., Concepcion A.R., Maus M., Shaw P., Portal-Celhay C., Zahra A., Kozhaya L., Weidinger C., Philips J., Unutmaz D., Feske S.* 2001. Ca^{2+} -signaling but not store-operated Ca^{2+} entry is required for the function of macrophages and dendritic cells. *J. Immunol.* 195 : 1202–1217.
- Wei M., Zhou Y., Sun A., Ma G., He L., Zhou L., Zhang S., Liu J., Zhang S.L., Gill D.L., Wang Y.* 2016. Molecular mechanisms underlying inhibition of STIM1–Orai1-mediated Ca^{2+} entry induced by 2-aminoethoxydiphenyl borate. *Pflugers Arch.* 468 : 2061–2074.
- Xie Q., Zhang Y., Zhai C., Bonanno J.A.* 2002. Calcium influx factor from cytochrome P-450 metabolism and secretion-like coupling mechanisms for capacitative calcium entry in corneal endothelial cells. *J. Biol. Chem.* 277 : 16559–16566.
- Xu X., Ali Sh., Li Y., Yu H., Zhang M., Lu J., Xu T.* 2016. 2-aminoethoxydiphenyl borate potentiates CRAC current by directly dilating the pore of open Orai1. *Sci. Rep.* 6 : 29304.
- Yamashita M., Somasundaram A., Prakriya M.* 2011. Competitive modulation of Ca^{2+} release-activated Ca^{2+} channel gating by STIM1 and 2-aminoethoxydiphenyl borate. *J. Biol. Chem.* 286 : 9429–9442.

THE EFFECT OF 2-AMINOETHOXYPHENYL BORATE ON STORE-DEPENDENT Ca^{2+} ENTRY IN PERITONEAL MACROPHAGES

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Store-dependent Ca^{2+} -entry is a universal mechanism of regulated Ca^{2+} entry in eukaryotic cells. To elucidate the pharmacological characteristics of store-dependent Ca^{2+} entry in macrophages, the effect of 2-aminoethoxydiphenyl borate (2-APB) on store-dependent Ca^{2+} entry, induced by endoplasmic Ca^{2+} -ATPases inhibitors thapsigargin

and cyclopiazonic acid as well as disulfide-containing immunomodulators glutoxim and molixan, was investigated in rat peritoneal macrophages. Using Fura-2AM microfluorimetry we have shown that in rat peritoneal macrophages, as well as in other cell types, 2-APB modulates store-dependent Ca^{2+} entry in a dose-dependent manner. At concentration of 25 μM 2-APB potentiates Ca^{2+} entry, while at concentrations of 50 and 100 μM it effectively inhibits store-dependent Ca^{2+} entry in macrophages. The results additionally confirm that Ca^{2+} entry induced by glutoxim or molixan occurs via store-dependent mechanism.

Keywords: 2-aminoethoxydiphenyl borate, peritoneal macrophages, store-dependent Ca^{2+} entry