# David Don Busath

January 8, 2019

Position:	Professor	Date of Birth:	August 5, 1952
		Place of Birth:	Salt Lake City, Utah
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# Academic Record:

University of Utah, Salt Lake City, UT	B.A. Physics & Honors (Cum laude)	1974
University of Utah, Salt Lake City, UT	M.D.	1978

# **Academic Positions:**

2011-2013	Adjunct Professor Research, Touro University Nevada
2004 - present	Professor, Department of Physiology and Developmental Biology, Brigham
-	Young University, Provo, Utah
1999 - present	Professor, Center for Neuroscience, Brigham Young University, Provo, UT
1996 - 2004	Professor, Zoology Department, Brigham Young University, Provo, UT
1995 - 1996	Associate Professor, Zoology Department, Brigham Young University,
	Provo, UT
1989 - 1995	Associate Professor with Tenure, Department of Physiology, Brown
	University, Providence, RI
1983-1989	Assistant Professor, Section of Physiology and Biophysics
	Brown University, Providence, RI
1980-1983	Research Associate, Department of Physiology and Biophysics
	University of Texas Medical Branch, Galveston, TX
1978-1980	Postdoctoral Fellow, Department of Physiology
	University of Rochester Medical School, Rochester, NY

#### **Honors and Awards:**

Markey Fellow, Mount Dessert Island Biological Laboratory, 1985 NIH Research Career Development Award, 1986-1991 Alcuin Fellowship for General Education Teaching (BYU), 2001-2003 Excellence in Research Award, College of Biology and Agriculture (BYU), 2002 Thomas B. Martin Professorship, College of Biology and Agric. (BYU), 2003-2006 Distinguished Faculty Award, Dept. of PDBio (BYU), 2010 Outstanding Research Award, College of Life Sciences (BYU), 2014

#### **Editorial Positions:**

Editor in Chief, Molecular Modeling and Computational Chemistry Review. 1997 – 2007 Associate Editor, Biochimica et Biophysica Acta, 1999-2004 Editorial Board, The Open Journal of Structural Biology, Bentham Press, 10/2007-2012 Editorial Board Member, Journal of Membrane Biology 2008-2015

#### Articles in journals (yellow: undergrads; cyan: graduate students):

- 1. Busath, D.D. and R.O. Stenerson. 1971. 2-particle, 3-particle and 4-particle spatial correlations among tertiary cosmic ray muons. Nuclear Physics B. 35:141-159.
- Eichwald, E.J., G. Pay, D. Busath, and C. Smith. 1976. Ischemic versus cytotoxic damage in white graft rejection - its relationship to hyperacute kidney rejection. Transplantation 22:86-93.
- 3. Begenisich, T. and D. Busath. 1981. Sodium flux ratio in voltage-clamped squid giant axons. Journal of General Physiology 77:489-502.
- 4. Busath, D. and G. Szabo. 1981. Gramicidin forms multi-state rectifying channels. Nature 294:371-373.
- 5. Prasad, K.U., T.L. Trapane, D. Busath, G. Szabo and D.W. Urry. 1982. Synthesis and characterization of 1-13 C-D-Leu 12, 14 Gramicidin A. International Journal of Peptide and Protein Research 19:162-171.
- 6. Busath, D. and T. Begenisich. 1982. Unidirectional sodium and potassium fluxes through the sodium channel of squid giant axons. Biophysical Journal 40:41-50.
- Prasad, K.U., T.L. Trapane, D. Busath, G. Szabo and D.W. Urry. 1982. Solid phase (Fmoc) synthesis and characterization of [1-13C Phe11]-Gramicidin B. J. Protein Chem. 1:191-202.
- Prasad, K.U., T.L. Trapane, D. Busath, G. Szabo and D.W. Urry. 1983. Synthesis and characterization of (1-13C) Phe9 Gramicidin A:effects of side chain variations. International Journal of Peptide and Protein Research 22:341-347.
- 9. Busath, D.D. and R.C. Waldbillig. 1983. Photolysis of gramicidin A channels in lipid bilayers. Biochimica et Biophysica Act. 736:28-38.

- 10. Busath, D. and G. Szabo. 1984. Atypical gramicidin A channels have increased field strength at one binding site. Biophysical Journal 45:75-76.
- Chabala, L.D., R.S. Morello, D. Busath, M. Danko, C.J. Smith, and T. Begenisich. 1986. Capture, transport, and maintenance of live squid (Loligo pealei) for electrophysiological studies. Pfluger Arch. 407:105-108.
- Jones, D., Hayon, E. and D. Busath. 1986. Tryptophan photolysis is responsible for gramicidin-channel inactivation by ultraviolet light. Biochimica et Biophysica Act. 861:62-66.
- 13. Busath, D.D., O.S. Andersen, and R.E. Koeppe II. 1987. On the conductance heterogeneity in membrane channels formed by gramicidin A. A cooperative study. Biophysical Journal 51:79-88.
- 14. Busath, D. and G. Szabo. 1988. Low conductance gramicidin A channels are head-tohead dimers of  $\beta$ 6.3-Helices. Biophysical Journal 53:689-695.
- 15. Busath, D. and G. Szabo. 1988. Permeation characteristics of gramicidin conformers. Biophysical Journal 53:697-707.
- Busath, D.D. and E. Hayon. 1988. Ultraviolet flash photolysis of gramicidin-doped lipid bilayers. Biochim. Biophy. Acta 944:73-78.
- Roeske, R.W., Hrinyo-Pavlin, T.P. Pottorf, R.S., Bridal, T., Jin, X., and D. Busath. 1989. Synthesis and channel properties of Tau-16 gramicidin A. Biochimica et Biophysica Acta 982:223-227.
- 18. Hemsley, G. and D. Busath. 1991. Small iminium ions block gramicidin channels in lipid bilayers. Biophysical Journal 59:901-908.
- Turano, B., Pear, M., and D. Busath. 1992. Gramicidin channel selectivity: molecular mechanics calculations for formamidinium, guanidinium, and acetamidinium. Biophysical Journal 63:152-161.
- 20. Bridal, T. and D. Busath. 1992. Inhibition of gramicidin channel activity by local anesthetics. Biochimica et Biophysica Acta. Biochim. Biophys. Acta 1107:31-38.
- 21. Bogusz, S., Boxer, A., and D.D. Busath. 1992. An SS1-SS2 β-barrel structure for the voltage-activated potassium channel. Protein Engineering 5:285-293.
- 22. Bogusz, S. and D.D. Busath. 1992. Is a  $\beta$ -barrel comprised of SS1 and SS2 from the voltage-gated potassium channel energetically feasible? Biophysical Journal 62:19-21.
- 23. Seoh, S. -A. and D. Busath. 1993. The permeation properties of small organic cations in gramicidin A channels. Biophysical Journal 64:1017-1028.
- Seoh, S.-A. and D.D. Busath. 1993. Formamidinium-induced dimer stablization and flicker block behavior in homo- and heterodimer channels formed by gramicidin A and Nacetyl gramicidin A. Biophysical Journal 65:1817-1827.
- 25. Seoh, S. and D. D. Busath. 1995. Gramicidin tryptophans mediate formamidiniuminduced channel stabilization. Biophysical Journal 68:2271-2279.

- 26. Hao, Y., Pear, M.R. and D.D. Busath. 1997. Molecular dynamics study of free energy profiles for organic cations in gramicidin A channels. Biophysical Journal 73: 1699-1716.
- Busath, D.D., Thulin, C.D., Hendershot, R.W., Phillips, L.R., Maughan, P., Cole, C.D., Bingham, N.C., Morrison, S., Baird, L.C., Hendershot, R.J., Cotten, M., and Cross, T.A. 1998. Noncontact dipole effects on channel permeation. I. Experiments with (5F-Indole)Trp13 gramicidin A channels. Biophysical Journal 75:2830-2844.
- Dorigo, A.E., Anderson, D.G., and Busath, D.D. 1999. Noncontact dipole effects on channel permeation. II. Trp conformations and dipole potentials in gramicidin A. Biophysical Journal 76: 1897-1908.
- 29. Andersen, O.S., H.J. Appell, E.Bamberg, D.D. Busath, R.E. Koeppe II, F.J. Sigworth, G. Szabo, D.W. Urry, and A. Woolley. 1999. Gramicidin channel controversy -- The structure in a lipid environment. Nature Structural Biology 6:609-611.
- Cotten, M., C. Tian, D.D. Busath, R.B. Shirts, and T.A. Cross. 1999. Modulating Dipoles for Structure-Function Correlations in the Gramicidin A Channel. Biochemistry 38:9185-9197.
- Fairbanks, T.G., C.L. Andrus, and D.D. Busath. 1999. Lorentzian noise in single gramicidin A channel formarmidinium currents. In. Gramicidin and Related Ion Channel-Forming Peptides. Novartis Foundation Symposium 225. John Wiley & Sons, Ltd. Chichester, UK. pp 74-92.
- Phillips, L.R., C.D. Cole, R.J. Hendershot, M. Cotton, T.A. Cross, and D.D. Busath. 1999. Noncontact dipole effects on channel permeation. III. Anomalous proton conductance effects in gramicidin. Biophysical Journal 77:2492-2501.
- Hollerbach, U., D.P. Chen, D.D. Busath, and B. Eisenberg. 2000. Predicting function from structure using the Poisson-Nernst-Planck equations: sodium current in the gramicidin A channel. Langmuir 16:5509-5514.
- Boda, D., D.D. Busath, D. Henderson, and S. Sokolowski.2000. Monte Carlo simulations of the mechanism for channel selectivity: The competition between volume exclusion and charge neutrality. J. Phys. Chem. B. 104:8903-8910.
- Thompson, N., G. Thompson, C.D. Cole, M. Cotton, T.A. Cross, and D.D. Busath. 2001. Non-contact dipole effects on channel permeation. IV. Kinetic model of 5F-Trp13 gramicidin A currents. Biophys. J. 81: 1245-1254.
- Anderson, D.G., R.B. Shirts, T.A. Cross, and D.D. Busath. 2001. Non-contact dipole effects on channel permeation. V. Computed potentials for fluorinated gramicidin. Biophys. J. 81: 1255-1264.
- Crozier, P.S., R.L. Rowley, N.B. Holladay, D. Henderson, and D.D. Busath. 2001. Molecular dynamics simulation of continuous current flow through a model biological membrane channel. Physics Review Letters 86:2467-2470.
- Crozier, P.S., D. Henderson, R.L. Rowley, and D.D. Busath. 2001. Model channel ion currents in NaCl-Extended Simple Point Charge Water solution with applied-field molecular dynamics. Biophysical Journal 81:3077-3089.

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- Boda, D., D. Henderson, and D.D. Busath. 2001. Monte Carlo study of the effect of ion and channel size on the selectivity of a model calcium channel. J. Physical Chemistry B 105:11574-11577.
- 40. Henderson, D., D.D. Busath, and R. Rowley. 2001. Fluids near surfaces and in pores and membrane channels. Progress in Surface Science 68:279-295.
- Markham, J.C., J.A. Gowen, T.A. Cross, and D.D. Busath. 2001. Comparison of gramicidin A and gramicidin M channel conductance dispersion properties. Biochimica et Biophysical Acta 1513:185-192.
- 42. Boda, D., D. Henderson, and D.D. Busath. 2001. Monte Carlo study of the effect of ion and channel size on the selectivity of a model calcium channel. J. Physical Chemistry B 105:11574-11577.
- Cole, C. D., A. S. Frost, N. Thompson, M. Cotten, T. A. Cross, and D. D. Busath. 2002. Non-contact dipole effects on channel permeation. VI. 5F- and 6F-Trp gramicidin channel currents. Biophys J. 83:1974-1986.
- 44. Ramakrishnan, V. and D. D. Busath. 2002. An inverting basket model for Band 3 obligate exchange. Journal of Theoretical Biology 215:215-226.
- 45. Gowen, J.A. J.C. Markham, S.E. Morrison, D.D. Busath, T.A. Cross, E.J. Mapes, and M.F. Schumaker. 2002. The role of Trp side chains in tuning single proton conduction through gramicidin channels. Biophysical Journal 83:880-898.
- 46. Yang, Y., D. Henderson, P. Crozier, R.L. Rowley, and D.D. Busath. 2002. Permeation of ions through a model biological channel: Effect of periodic boundary conditions and cell size. Molecular Physics. 100:3011-3019.
- 47. Boda, D., D. Henderson, and D.D. Busath. 2002. Monte Carlo study of the selectivity of calcium channels: Improved geometrical model. Molecular Physics. 100:2361-2368.
- 48. Boda, D., D.D. Busath, B. Eisenberg, D. Henderson, and W. Nonner. 2002. Monte Carlo simulations of selectivity in neuronal Na channels: Charge space competition. Phys. Chem. Chem. Phys. 4, 5154-5160.
- 49. Fu, F.-N. D.D. Busath, and B.R. Singh. 2002. Spectroscopic analysis of low pH and lipid induced structural changes in type A botulinum neurotoxin relevant to membrane channel formation and translocation. Biophys Chem. 99:17.
- 50. Duffin, R.L., Garrett, M.P., Flake, K.B., Durrant, J.D., and D. D. Busath. 2003. Modulation of lipid bilayer interfacial dipole potential by phloretin, RH421, and 6-ketocholestanol as probed by gramicidin channel conductance. Langmuir 19:1439-1442.
- 51. Yang, Y., D. Henderson, and D.D. Busath. 2003. Applied-field molecular dynamics study of a model calcium channel selectivity filter. J. Chem. Phys. 118: 4213-4220.
- Bingham, N.C., N.E.C. Smith, T.A. Cross, and D.D. Busath. 2003. Molecular dynamics simulations of Trp side-chain conformational flexibility in the gramicidin A channel. Biopolymers 71: 593-600.

- 53. Yang, Y., D. Henderson, and D.D. Busath. 2004. Calcium block of sodium current in a model calcium channel: Cylindrical atomistic pore with glutamate side chains. Molecular Simulation 30:77-80.
- 54. Yang, Y., M. Berrondo, D. Henderson, and D. Busath. 2004. The importance of water molecules in ion channel simulations. J. Phys.: Condens. Matter 16: S2145-S2148.
- 55. Hughes T., B. Strongin, F.P. Gao, V. Vijayvergiya, D.D. Busath, and R.C. Davis. 2004. AFM visualization of mobile influenza A M2 molecules in planar bilayers. Biophysical Journal 87: 311-22.
- 56. Ramakrishnan, V., D. Henderson, and Busath, D.D. 2004. Applied field nonequilibrium molecular dynamics simulations of ion exit from a beta-barrel model of the L-type calcium channel. Biochim Biophys Acta 1664: 1-8.
- Vijayvergiya V., R. Wilson, A. Chorak, P.F. Gao, T.A. Cross, and D.D. Busath. 2004. Proton conductance of influenza virus M2 protein in planar lipid bilayers. Biophysical Journal 87: 1697-1704.
- Hu, J., R. Fu, K. Nishimura, L. Zhang, H.-X. Zhou, D. D. Busath, V. Vijayvergiya and T.A. Cross. 2006. Histidines: Heart of the H+ channel from influenza A virus. Proceedings of the National Academy of Sciences 103:6865-6870.
- Durrant, J.D., D. Caywood, D.D. Busath. 2006. Tryptophan contributions to the empirical free-energy profile in gramicidin A/M heterodimer channels. Biophysical Journal 91: 3230-3241.
- Moffatt, C., V. Vijayvergiya, P.F. Gao, T.A. Cross, D.J. Woodbury, and D.D. Busath. 2008. Proton transport through influenza A virus M2 protein reconstituted in vesicles. Biophysical Journal 94:434-445.
- 61. Miller, C.E., D.D. Busath, B. Strongin, and J. Majewski. 2008. Integration of Ganglioside GT<sub>1b</sub> Receptor into DPPE and DPPC Phospholipid Monolayers: An X-ray Reflectivity and Grazing Incidence Diffraction Study. Biophysical Journal 95:3278-3286.
- 62. Mustafa, M. and D.D. Busath. 2009. The gramicidin channel ion permeation free-energy profile: direct and indirect effects of CHARMM force field improvements. Interdisciplinary Science: Computational Life Sciences 1:113-127.
- 63. Mustafa M., D.J. Henderson, and D.D. Busath. 2009. Free-energy profiles for ions in the influenza M<sub>2</sub>-TMD channel. Proteins 76:794-807.
- Mustafa M., D.J. Henderson, and D.D. Busath. 2009. Computational studies of gramicidin permeation: An entryway sulfonate enhances cation occupancy at entry sites. Biochim. Biophys. Acta. 1788:1404-1412.
- 65. Spohr, E., E. Sovyak, A. Trokhymchuk, and D.D. Busath. 2009. Electrostatic control of occupancy and valence selectivity in a charged nanometer-sized cylindrical pore. Materialwissenschaft und Werkstofftechnik 40: 247-254.
- 66. Jones, T.L., R. Fu, F. Nielson, T.A. Cross, and D.D. Busath. 2010. Gramicidin channels are internally gated. Biophys. J. 98:1486-1493.

- 67. Sharma, M., M. Yi, D. Hao, H. Qin, E. Peterson, D. D. Busath, H.-X. Zhou, and T. A. Cross. 2010. Insights into the mechanism of the influenza A proton channel from a structure in a lipid bilayer. Science 330:509-512.
- Peterson, E., T. Ryser, S. Funk, D. Inouye, M. Sharma, H. Qin, T. A. Cross, and D. D. Busath, Functional reconstitution of influenza A M2(22-62). 2011. Biochim. et Biophys. Acta, Biomembranes 1808:516-521 (Published electronically 2010).
- Sharma M, C. Li, D. D. Busath, H.-X. Zhou, and T. A. Cross. 2011. Drug sensitivity, drugresistant mutations, and structures of three conductance domains of viral porins. Biochim. et Biophys. Acta., Biomembranes 1808:538-546.
- Mellor, B., E. Cruz Cortés, D. D. Busath, and B. Mazzeo. 2011. Method for estimating the internal permittivity of proteins using dielectric spectroscopy. J. Phys. Chem. B. 115:2205-2213.
- 71. Mellor, B. L., S. Khadka, D. D. Busath, and B. Mazzeo. 2011. Influence of pKa shifts on the calculated dipole moments of proteins. Protein Journal 30:490-498.
- 72. Nelson, S. C., S. K. Neeley, E. D. Melonakos, J. D. Bell, and D. D. Busath. 2012 Fluorescence anisotropy of diphenylhexatriene and its cationic trimethylamino derivative in liquid dipalmitoylphosphatidylcholine liposomes: Opposing responses to isoflurane. BMC Biophysics 5:5.
- 73. Cross, T. A., H. Dong, M. Sharma, D. D. Busath, H.-X. Zhou. 2012. M2 Protein from Influenza A: From multiple structures to biophysical and functional insights. Current Opinions Virology 2:128-133.
- 74. Mazzeo, B. A. and D. D. Busath. 2013. From molecular dynamics to fluorescence anisotropy of fluorophores bound to oriented structures. J. Comp. Phys. 232:482-497.
- 75. Shin, S. I., Andersen, D. J., Hansen, D. M., Yorgason, J. T., Schilaty, N. D., Busath, D. D., and S. C. Steffensen. 2013. Connexin-36 knock-out mice have increased threshold for kindled seizures: Role of GABA inhibition. Biochem & Pharmacol S1: 006. doi:10.4172/2167-0501.S1-006.
- Kolocouris, A., Tzitzoglaki, C., Johnson, B., Zell, R., Wright, A., Cross, T.A., Tietjen, I., Fedida, D. & D. Busath. 2014. Adamantanes with persistent in vitro efficacy against H1N1 (2009) Influenza A. J. Med. Chem. 57:4629-4639.
- 77. Gleed, M. and D. D. Busath. 2015. Why bound amantadine fails to inhibit proton conductance according to simulations of the drug-resistant influenza A M2 (S31N). J. Phys. Chem. B 119 (3):1225-31.
- Durrant, M.G., Eggett, D.L., and D.D. Busath. 2015. Investigation of a recent rise of dual amantadine-resistance mutations in the Influenza A M2 Sequence. BMC Genetics 16 (Suppl. 2): S3.
- Gleed, M. L., Ioannidis, H., Kolocouris, A., and D.D. Busath. 2015. Resistance-mutation (N31) effects on drug orientation and channel hydration in amantadine-bound influenza A M2. J. Phys. Chem. B 119: 11548-11559.

- Gordon, N.A., McGuire, K.L., Wallentine, S.K., Mohl, G.A., Lynch, J.D., Harrison, R.G., Busath, D.D. 2017. Divalent copper complexes as influenza A M2 inhibitors, Antiviral Research 147: 100-106, doi: 10.1016/j.antiviral.2017.10.009.
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- Mohl, G., Liddle, N., Nygaard, J., Dorius, A., Lyons, N., Hodek, J., Weber. J., Michaelis, D., Busath, D.D. Novel Influenza Inhibitors Designed to Target PB1 Interactions with Host Importin RanBP5. Antiviral Research, In Press.

#### **Opinions/Editorials**

- 1. Andersen O. S., H. J. Apell, E. Bamberg, D. D. Busath, R. E. Koeppe 2nd, F. J. Sigworth, G. Szabo, D. W. Urry, and A. Woolley. 1999. Gramicidin channel controversy--the structure in a lipid environment. Nat Struct Biol. 6:609; discussion 611-2.
- 2. Separovic, F., J. A. Killian, M. Cotten, D. D. Busath, and T. A. Cross. 2011. Modeling the membrane environment for membrane proteins. Biophysical Journal 100:2073-2074.
- 3. Busath D. D., D. J. Woodbury, and A. Frost. 2012. Endosis and exosis: new names for fusion and budding. J Membr Biol. 245:759-760.

#### Chapters in Books, Invited Reviews, Proceedings, Etc.:

- Szabo, G. and D.D. Busath. 1983. Ion movement through membrane channels. In: "Membrane Biophysics: Physical Methods in the Study of Biophysical Systems." M.A. Dinno, A.B. Callahan, T.C. Rosell, Eds. Alan R. Liss, Inc., New York.
- Busath, D., Hemsley, G., Bridal, T., Pear, M., Gaffney, K. and M. Karplus. 1988. Guanidinium as a probe of the gramicidin channel interior. In: "Transport through Membranes: Carriers, Channels and Pumps." A. Pullman, J.J. Jortner, B. Pullman Eds. Kluwer Academic Publishers, Boston (Norwell, MA 02061).
- Chen, I. and D. Busath. 1990. Animating a cellular transport mechanism. Pixel 1:16-23.
- Busath, D.D. 1993. The use of physical methods in determining gramicidin channel structure and function. Annual Reviews of Physiology 55:473-501.
- Henderson, D., D. D. Busath, R. L. Rowley, P. S. Crozier, and D. Boda. 2001. Simulation study of channels in biological membranes. Proceedings of the International Conference on Computational Nanoscience. Pp. 45-48.
- Yang, Y., Boda, D., Henderson, D. and D. Busath. 2002. Computer simulation studies of the selectivity and conductance of a model calcium channel. Journal of Computational Electronics. 1: 353-357.
- Caywood, D. and D.D. Busath. 2002. Oriented Gramicidin M Gramicidin A heterodimers: Rectification decreases with increased ion concentration. In: "Membrane Interacting Peptides and Proteins 2002." F. Heitz, ed., Research Signpost, Kerala, India. Pp 147-153.
- Boda, D., Busath, D.D., and Henderson. 2002. Simulation of the selectivity of a calcium channel.

Applied Surface Science 196:154-156.

- Yang, Y., D. Henderson, and D.D. Busath. 2004. Calcium block of sodium current in a model calcium channel: Cylindrical atomistic pore with glutamate side chains. Molecular Simulations 30:75-80.
- Boda D., T. Varga, D. Henderson, D.D. Busath, W. Nonner, D. Gillespie, and B. Eisenberg. 2004. Monte Carlo simulation study of a system with a dielectric boundary: Application to ion channel selectivity. Molecular Simulations 30:89-96.
- Busath, D., D. Henderson, and S. Sokolowski. 2004. Density functional theory for an electrolyte in a cylinder: The selectivity of a calcium channel. J. Phys.: Condens. Matter 16: S2193-S2201.
- Busath, D. D. 2009. Influenza A M2: Channel or Transporter? In "Advances in Planar Lipid Bilayers and Liposomes," A. Leitmannova Liu and Aleš Iglič, editors, Burlington: Academic Press,10:161-201.

#### **Grants Awarded:**

9/1/78 - 8/31/80	NIH NRSA 5F32 NS-06084-02 Postdoctoral Fellowship
	The sodium flux ratio in squid axon sodium channels.
	Sponsor: Ted Begenisich. \$26,300.
7/1/83 - 6/30/85	American Heart Association, Texas Affiliate
	The pore structure in transmembrane channels.
	P.I. David Busath. \$53,906, direct costs.
8/1/83 - 7/31/86	NIH 1 RO1 GM33361
	Molecular interactions inside a transmembrane pore.
	P.I. David Busath. \$166,088, direct costs.
6/1/85 - 7/31/85	Mount Desert Island Biol. Lab. Markey Fellowship.
	The photolysis of transmembrane channels in nerve and oocyte.
	P.I. David Busath. \$5,000, direct costs.
7/1/86 - 6/30/91	NIH 1 K04 NS01085 Research Career Development Award
	Gramicidin channel blockade induced by guanidinium.
	P.I. David Busath. \$261,198, direct costs.
8/1/86 - 7/31/91	NIH R01 GM33361
	Molecular interactions inside a transmembrane pore.
	P.I. David Busath. \$421,517, direct costs.
4/1/87 - 3/31/88	NIH BRS Shared Instrumentation Grant
	Peptide Synthesizer and Peptide Sequencer.
	P.I. Wayne Bowen. \$101,000, direct costs.
12/1/88 - 11/30/89	NIH DRR BRS Shared Instrumentation Grant.
	Molecular Modeling Network.
	P.I. John W. Suggs. \$81,080, direct costs.
3/1/95 - 2/28/00	NIH RO1 AI 23007
	Correlations: Structures-Dynamics-Functions in Gramicidin
	P.I. Timothy A. Cross; Sub-contract P.I. David Busath.
	\$1,067,657 total costs.
	\$213,215, total costs in the Brown/BYU subcontract.
10/1/96-9/30/99	NSF Academic Research Infrastructure. Acquisition of a Computational

	Chemistry Server. P.I. Randall B. Shirts; Co-P.I. David Busath and others.
	\$100,000 direct costs. (Matched by BYU matching funds)
3/1/00-2/29/05	NIH RO1 AI 23007
	Correlations: Structure-Dynamics-Functions in Channels.
	P.I. Timothy A. Cross; Sub-contract P.I. David Busath.
	\$1,771,739 total costs. \$462,353 total costs in the BYU subcontract.
4/1/00-3/31/03	NSF Research Experience for Undergraduates (REU Site)
	"Neuroscience Research Experiences for Undergraduates Site at BYU"
	P.I. Edwin Lephart, \$237,829 total costs.
3/1/01-12/31/01	BYU "Neuroscience Environment for Mentoring" P.I. Edwin Lephart,
	\$34,650.00 total costs.
3/1/05-2/29/10	NIH RO1 AI 23007
	Correlations: Structure-Dynamics-Functions in Channels.
	P.I. Timothy A. Cross; Sub-contract P.I. David Busath.
	\$1,805,550 total costs. \$353,827 total costs in the BYU subcontract.
1/1/08-9/1/08	Botulinum Toxin Research Associates
	Subcutaneous Botox Effects on Electrically Induced Seizures.
	P.I. David Busath; co-P.I. Scott Steffensen
	\$16,000 total costs.
8/1/11-7/31/15	NIH RO1 AI 23007
	Correlations: Structure-Dynamics-Functions in Channels.
	P.I. Timothy A. Cross; Sub-contract P.I. David Busath.
	~\$500,000 total costs in the BYU subcontract.

# Internal funding:

Mentoring Environment Grants from ORCA, BYU of \$20,000 in 2004, 2006, 2008, 2011, 2012, and 2014
Robert Gardner Fund from the College of Life Sciences, BYU of \$30,000 in 2009 to search for mechanisms of food intelerance
Seed award of \$10,000 from the Fulton College of Engineering, BYU to Brian Mazzeo and me for "Dielectric Spectroscory Assessment of Influenze A M2 Drug Din ding?" in 2010
Capital equipment award from Depts. of PDBio and Chem/Biochem, BYU of \$90,000 to Dixon
Woodbury and me for a circular dichroism spectrometer in 2010
Tech Transfer Bridging Award of \$25,000 for development of influenza drugs in 2012
Capital equipment award from Dept. of PDBio, BYU of \$18,500 to John Bell and me for a fluorescence lifetime spectrophotometer in 2013.
Student Entrepreneur Mentoring Award from the BYU Rollins Center of \$5,000 in 2014.
CHIRP Award from Dept. of Chemistry and Biochemistry, BYU of \$20,000 to spark translational research to Roger Harrison and me in 2014.
Research award from the Magnetic Resonance Imaging Facility, BYU of \$1,000 to help initiate an
fMRI study of chronic pain in 2014; additional \$9,000 in 2015.
Gift from the Sorenson Legacy Foundation, 2016, \$25,000 for "Identify new methods to block neuropathic pain."
College of Life Sciences Technology Transfer Award, 2016, \$15,000 for "Leveraging Three Anti-

Influenza Drug Patent Applications By Development of a Zebrafish Larvae Virus-Injection Assay." Arminda Suli, Co-PI.

College of Life Sciences Technology Transfer Award, 2017, \$15,000 for "Continued Development of a Zebrafish Larvae Virus-Injection Assay." Arminda Suli, Co-PI.

#### **Provisional Patent Applications Submitted:**

- 9/1/2017 App No. 62/553,732 "Novel Influenza Chemotherapeutics: Nuclear Import Inhibitors of the Influenza Polymerase," Inventors: David D. Busath, David Michaelis, and Greg Mohl.
- 4/5/2017 App No. 62/481,909 "Novel Influenza Chemotherapeutics: Nuclear Import Inhibitors of the Influenza Polymerase," Inventors: David D. Busath, Greg Mohl, David Michaelis
- 1/30/2017 App No. 62/451,988 "Novel Influenza Chemotherapeutics: Nuclear Import Inhibitors of the Influenza Polymerase," Inventors: Greg Mohl and David D. Busath.
- 9/23/2014 App No. Not filed. "Antiviral-Metal Complexes for Treatment of Influenza Virus Infections," Inventors: David D. Busath, Timothy Cross, David Fedida, Nathan Gordon, Roger G. Harrison, Antonios Kolocouris, Michaela Schmidtke, Roland Zell.
- 6/12/2014 App No. 61/997,888 "Divalent Copper Compounds as Inhibitory Agents of Influenza A," Inventors: Nathan Gordon, Roger Harrison, David D. Busath.
- 12/23/2013 App No. 61/920,359 "A Novel Set of Antiviral Drugs for M2/S31N-bearing Influenza A. Inventors: David D. Busath, F. Brent Johnson, Antonios Kolocouris.
- 2/2/2013 App No. 61/760,060 "A Novel Set of Antiviral Drugs for M2/S31N-bearing Influenza A," Inventors: David D. Busath, F. Brent Johnson, Antonios Kolocouris.
- 9/29/2011 App No. 61/626,618 "A Novel Set of Antiviral Drugs for M2/S31N-bearing Influenza A. Inventors: David D. Busath, F. Brent Johnson, Antonios Kolocouris.

## Patent Cooperation Treaty (PCT) Patent Applications Submitted:

- 6/12/2015 App No. PCT/US15/35604 "ORGANO-TRANSITION METAL COMPLEXES FOR THE TREATMENT OF VIRAL INFECTIONS," Inventors: Nathan Gordon, Kelly McGuire, Spencer Wallentine, James Clark, Roger Harrison, David D. Busath
- 2/2/2014 App No. PCT/US14/14359 "A Novel Set of Antiviral Drugs for M2/S31Nbearing Influenza A." Inventors: David D. Busath, F. Brent Johnson, Antonios Kolocouris.

## **Full Patent Applications Submitted:**

12/12/2016 App No. 15/318,198 "ORGANO-TRANSITION METAL COMPLEXES FOR

THE TREATMENT OF VIRAL INFECTIONS," Inventors: Nathan Gordon, Kelly McGuire, Spencer Wallentine, James Clark, Roger Harrison, David D. Busath (according to Faculty Profile/TTO).

8/2/2015+ App No. 14/765,539 "Antiviral Compounds." Inventors: David D. Busath, F. Brent Johnson, Antonios Kolocouris. (USPO). Also submitted within one month to patent offices in Korea, EP, AU, and Japan.

#### **Patents Awarded:**

12/12/2017 USPO #9,840,465 "Antiviral Compounds." Inventors: David D. Busath, F. Brent Johnson, Antonios Kolocouris.

# **Invited Lectures:**

9/1985	"Guanidinium Blocks in Gramicidin Channels". Department of Physiology, Emory
	University, Atlanta, GA.
11/1985	"Iminium Ion Blocks in Gramicidin Channels". Department of Physiology, Yale
	University, New Haven, CT.
3/1986	"Small iminium ions block gramicidin channels in lipid bilayers". Department of
	Pharmacology and Toxicology. Dartmouth Medical School. Hanover, NH.
5/1988	"Guanidinium as a probe of the gramicidin channel interior". Jerusalem
	Symposium on Quantum Chemistry and Biochemistry, Jerusalem, Israel.
5/1988	"Guanidinium as a probe of gramicidin channels". Laboratoire de Physicochimie
	des Systemes Polyphases. C.N.R.S. Montpellier, France.
9/1990	"Guanidinium as a probe of gramicidin channel structure". Department of
	Physiology, University of Illinois, Champaign, IL.
2/1992	"The voltage-gated K+ channel: a beta-barrel structure". Section of Physiology,
	Brown University, Providence, RI.
2/28/1992	"Modeling the pore of voltage-gated K+ channels". Department of Biophysical
	Sciences, SUNY, Buffalo, NY.
2/27/1992	"Molecular modeling of the voltage-gated K+ channel". Department of Physiology
	and Biophysics, University of Rochester, Rochester, NY.
3/26/1992	"The structure of voltage gated potassium channels". Physics Dept., Boston
	University, Boston, MA.
5/16/1992	"An SS1-SS2 beta barrel model for the voltage-gated K+ channel". Department of
	Physiology, Albert Einstein Medical School, Bronx, NY.
11/5/1992	"Molecular modeling of large biomolecules". Department of Chemistry, University
	of Massachusetts at Dartmouth, Dartmouth, MA.
3/19/1993	"Molecular modeling of voltage-gated potassium channels". Department of
	Chemistry, State University of New York at Binghamton, Binghamton, NY.
4/1993	"Molecular modeling of gramicidin and potassium channels". Department of
	Chemistry, University of Montreal at Montreal, Quebec, Canada.
5/1993	"Molecular modeling of the voltage-gated K+ channel". Pfizer Pharmaceutical
	Corporation, Groton, CT.

4/1994	"Gramicidin transport and dynamics with iminium ions." Friends of the Membrane.
<b>a</b> /1 a a <b>f</b>	Department of Physiology. Cornell University Medical School, New York, NY.
2/1995	"Binding selectivity in Voltage-gated channels". Dept. Of Physiology, University
	of Utah Medical School, Salt Lake City, Utah.
2/1995	"Binding selectivity in Voltage-gated channels". Zoology Dept, Brigham Young
	University, Provo, Utah.
3/1996	"Organic Cation Permeability of the Gramicidin Channel". Theoretical Physics
	Group, BYU, Provo, Utah.
9/1996	"Organic Cation Permeability of the Gramicidin Channel". Zoology Dept., BYU,
	Provo, Utah.
10/1996	"Organic Cation Permeability of the Gramicidin Channel". Microbiology Dept.,
	BYU, Provo, Utah.
11/18/98	"Lorentzian noise in single gramicidin A channel formamidinium currents."
	Novartis Symposium. London, UK.
6/98	"Formamidinium noise in gramicidin channels." Dept. of Physiology and
	Biophysics. Rush Medical School. Chicago, IL
9/15/99	"Proton permeation in gramicidin analogs: An assay of water reorientation rates."
	Satellite Symposium of XIII Int. Biophysics Congress. Center for Cellular &
0.00	Molecular Biology. Hyderabad, India.
9/24/99	"The influence of polar side chains on channel conductance: Proton conductance in
	gramicidin." MBBG Seminar. University of Delhi, South Campus. New Delhi,
1/12/00	India "The big base of a second balancies," New linear Network last Secondary Mathematics
1/13/00	Dent DVL Prove UT
2/2/00	"Gramiaidin a model ion channel" NSE Program Directors Seminar Arlington
2/2/00	$V\Delta$
2/8/00	"Using fluorination to explore the electric field in gramicidin channels." Dept. of
2/0/00	Anatomy and Neurobiology. University of Utah. Salt Lake City, UT.
2/10/00	"Using fluorination to explore the electric field in gramicidin channels." Center
	for Neuroscience. Brigham Young University, Provo, UT.
4/23/01	"Determinants of channel permeability." Dept. of Physiology and Biophysics. Rush
	University Medical School. Chicago, IL
4/25/01	"Proton Exit, Not Water-Reorientation, is Rate Limiting for Proton Currents in
	Gramicidin Channels." Dept. of Physiology. Loyola University Medical Center.
	Chicago, IL.
5/4/01	"Issues in ion transport by protein channels." BYU Workshop on Electrolytes and
	Interfaces. Dept. of Chemistry. Brigham Young University. Provo, UT.
12/5/01	"Excitability begins here: Voltage-gated channels in nerve and muscle." Physics
	Dept. Colloquium. Brigham Young University, Provo, UT.
3/02	"Excitable channels in nerve and muscle." Dept. of Chemistry and Biochemistry,
C 10 0	Brigham Young University, Provo, UT.
6/02	"The structure and function of voltage gated channels." Neuroscience REU Summer
7/02	Seminar, Brigham Young University, Provo, UT.
//02	"Unannel behaviors of the M2 protein from Influenza A virus." Dept. of Chemistry,
	FIORIDA State University, Talianassee, FL.

11/03	"M2 Protein Function and Images in Planar Bilayers." M2 Workshop, Northwestern University, Evanston, Illinois.
1/04	"M2 Protein Function and Images in Planar Bilayers." PDBio Departmental Seminar, Brigham Young University, Provo, UT.
10/5/2004	"Selectivity of Ion Channels." Beckmann Institute Lab Meeting. University of Illinois at Champaign-Urbana. Urbana, IL.
12/17/05	"Tryptophan Fluorination Effects on Gramicidin Channel Conductance." Fluorine NMR Applications Symposium. Pacifichem 2005. Honolulu, HI.
3/15/06	"An Achilles Tendon in the Influenza Virus: M2." Department of Biology, Utah Valley State College, Orem, UT.
11/06	"Progress report on Influenza M2 channel behavior." Cross Lab Group. Dept. of Chemistry and Biochemistry. NHMFL. Florida State Univ. Tallahassee, FL.
10/16/07	"Molecular Modeling of Lipid Bilayers and Channel Forming Peptides." Richard Rowley Lab Group. Dept. of Chemical Engineering. Brigham Young University. Provo, UT.
10/25/07	"A Proposal for Radical Change in NIH Funding." NIH Peer Review Consultation Meeting. San Francisco, CA.
8/24/07	"Biophysical Measurements with Botulinum/A Toxin", Dartmouth Botulinum Toxin Symposium, Dartmouth, MA
3/10/08	"Single Channel Studies With Influenza A M2 (C19,50S) From E. Coli Inclusion Bodies." Influenza Research Group, Northwestern University, Evanston, IL.
4/22/11	"Influenza A M2 is a Drug Target." Toto Olivera Lab Meeting Seminar, Dept. of Biology, University of Utah, Salt Lake City, UT.
10/13/11	"Can we Stop Pandemic Flu with an Anti-viral Drug?" BYU Dept. of Physiology and Developmental Biology Seminar, Provo, UT
11/17/11	"Can we Stop Pandemic Flu with an Anti-viral Drug?" BYU Current Topics in Molecular Life Sciences Seminar Series, Provo, UT
9/18/2012	"Can we Stop Pandemic Flu with an Anti-viral Drug?" Dept. of Physics, Solid State Group Seminar Series, Provo, UT
5/15/2013	"Persistent in vitro inhibition of influenza A by amantadine analogs." William DeGrado Research Group Meeting, San Francisco, CA
9/12/2013	"Persistent Blockers for Modern Influenza A." Seminar, BYU Dept. of Physiology and Developmental Biology, Provo, UT
2/17/2014	"Influenza A Blockers with Reduced Resistance Formation." Conference Platform Presentation, 58 <sup>th</sup> Annual Meeting, Biophysical Society, San Francisco, CA
2/08/2015	"Molecular dynamics of amantadine block in M2 of influenza A: WT vs S31N." Conference Platform Presentation, 59 <sup>th</sup> Annual Meeting, Biophysical Society, Baltimore, MD. Lecture presented by student, Mitchell Gleed.

# International Symposium Lectures:

5/1988	"Guanidinium as a probe of the gramicidin channel interior". Jerusalem
	Symposium on Quantum Chemistry and Biochemistry, Jerusalem, Israel.
11/18/98	"Lorentzian noise in single gramicidin A channel formamidinium currents."
	Novartis Symposium. London, UK.

	Dusatii 1
9/1999	"Proton permeation in gramicidin analogs: An assay of water reorientation rates."
	Satellite Symposium of XIII International Biophysics Congress on Membranes,
	Sensors, and Cell Surfaces. Hyderabad, India.
12/13/05	"Flourinated gramicidin channels: effects on the single channel conductance."
	Symposium, Pacifichem 2005. Honolulu, Hawaii.
6/23/11	"Influenza A M2 is a Drug Target." Ion Channel Workshop, Vancouver, BC,
	Canada.
6/26/2013	"Persistent in vitro inhibition of influenza A by amantadine analogs." Ion Channel
	Retreat, Vancouver, BC

# **Teaching Experience:**

1984-1991	Bio 110, Cell Physiology and Biophysics. Course director (1985).
1985-1989	Bio 117, Mammalian Physiology.
1992-1994	Bio 117, Mammalian Physiology. Course director (1993-1995).
1986-1994	Chem/Bio 121, Molecular Modeling. Course director (1986, 1988, 1990-1995).
1993-1995	Bio 80, Introduction to Physiology.
1995-present	Zool 260, Physiol. & Dev. Biol. 220, Introduction to Human Anatomy
1997-1999	Zool 562, Neurophysiology
2000-2003	Neuro 105, Introduction to Neuroscience
2000-2003	Chem 489, Structural Biochemistry
2000-2002	Neuro 480, Advanced Neuroscience
2000-2001	Neuro 480, Neuroscience Lab
2003-present	Zoo 460, Physiol. & Dev. Biol. 362, Human Physiology
2004-present	Physiol. & Dev. Biol. 568, Electrophysiology and Cellular Biophysics
2005-present	Physiol. & Dev. Biol. 550R, Molecular Dynamics Simulations Lab
2011-present	Physiol. & Dev. Biol. 450R, Biophysics Research Training

## Trainees:

# Postdoctoral Fellows:

9/1986-12/1988	Dr. Irina Vayl. Molecular definition of bilayer surface tension.
1/1991-1/1992	Dr. Andrea Dorigo. Electrostatic impact of Trp side chain in gramicidin channels, free energy profile of guanidinium transport in gramicidin
	channels.
9/1996-10/1997	Dr. Craig Thulin. Mechanism of Conductance increase by fluorination of Trp <sup>13</sup> in gramicidin A.
1/2002-1/2006	Dr. Viksita Vijayvergiya. Influenza M2 reconstitution and channel activity.

# Ph.D. Recipients:

1992	Xian-Zheng Jin.	Tau-16 gramicidii	n channel properties.
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- 1993 Sang Ah Seoh. Organic cation selectivity in gramicidin channels.
- 1994 Steve Bogusz. Molecular modeling of voltage-gated channels.

- 1995 Yili Hao. Free energy profile for transport of small amines in gramicidin channels.
- 2002 Vivek Ramakrishnan. Cation flow selection in model calcium channels.
- 2008 Morad Alawneh. (with Doug Henderson) Molecular dynamics of M2 and gramicidin. Current doctoral student: Kelly McGuire

#### Master's Degree Recipients:

- 1989 Terry Bridal. The inactivation of gramicidin channels by local anesthetics.
- 1999 Chad Cole. Fluorination effects on gramicidin channel permeation.
- 2004 Travis Hughes. Atomic force imaging of influenza M2 proteins.
- 2005 Jacob Durrant. Estimation of the tryptophan potential in gramicidin channels.
- 2005 Mario Pinoli. Stability and permeability of the polyglutamine µ-helix in simulations.
- 2007 Brad Strongin. Thermodynamic evidence that ganglioside-mediated insertion of Botulinum A into the cholinergic nerve ending may precede endocytosis and acidification: a Langmuir film study.
- 2011 Emily Peterson. Proteoliposome Proton Flux Assays Establish Net Conductance, pH-Sensitivity, and Functional Integrity of a Novel Truncate of the M2 Ion "Channel" of Influenza A.
- 2013 Nathan A. Gordon. Divalent Copper Compounds as Inhibitory Agents of Influenza A.

## **Intramural Service:**

Member BYU Internal Review Board 1999-2003

Member College Faculty Rank and Status Committee 2005-2010 Member Board of Directors, Magnetic Resonance Imaging Facility 2012-2014 Member, Departmental Faculty Rank and Status Committee 2009-2013 Chair, Departmental Faculty Rank and Status Committee 2013 - present

# **Extramural Service:**

NIH Small Business Grant Reviewer 11/86, 6/87 NIH Physiology Study Section (ad hoc members) 2/88 National President, Masscomp User's Society 4/88-10/89 Organizer of the Gramicidin Club and its Banquet (-30 members), which met annually since 1987. NSF Computers in Education Grant Reviewer 8/90 Biophysical Journal, Biophysica Biochimica Acta, Biochemistry, European Reviewer: Biophysics Journal, Journal of Biological Chemistry. NRC Howard Hughes Predoctoral Fellowship Panelist 2/95, 2/98, 2/00 NRC Associateship Program Panelist 2/96-2/99. Spearheaded Invitation of Biophysical Society to meet in Salt Lake City in 2006. Spearheaded Formation of Permeation/Transport Biophysics Subgroup in the Biophysical Society, Feb. 2002; president of the group 2003-2006. NIH Ad Hoc Reviewer 11/03 NSF Ad Hoc Reviewer 8/03 NIH Biophysics of Neural Systems Study Section (ad hoc member) 10/07

#### **Professional Society Memberships:**

Biophysical Society (1978 - present) New York Academy of Sciences (1984-1992) The American Association for the Advancement of Science (1985-1999) American Chemical Society (1985-1993; 2005 – 2013) The Society of General Physiologists (1986-1995) Boston Channel Group (1986-1995) The American Physiological Society (1987-1995) The Protein Society (1987-1993) Antiviral Society (2011 – present) International Society for Antiviral Research (2012 – present)