American Chemical Society

National Awards Nomination Packet

Alfred Bader Award in Bioinorganic or Bioorganic Chemistry:2018 for: R Britt

Received: 10/30/2015 Cycle Year: 2

"For pioneering pulse EPR spectroscopy characterizing the Photosystem II Oxygen Evolving Complex, and continuing EPR spectroscopy probing a wide variety of important metalloenzymes and catalysts"

NOMINATOR:

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Have you discussed this award nomination with the nominee?

Yes

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NOMINEE:

R Britt Tel: (916)795-0157 UC Davis Fax: (530)752-8995

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ACS Current Member: Yes Years of Service: 20

Date of birth: 01/01/1956

Present Position: Distinguished Professor of

Chemistry Academia

Industry: Academiá

• Does the nominee employ and require good safety protocols and practices in Yes

his/her laboratory?

What is the nominee's present position?
 Distinguished Professor of

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What professional discipline does the nominee work in?

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Alan L. Balch Distinguished Professor Former Chair of Chemistry, AAAS and ACS Fellow e-mail: albalch@ucdavis.edu

October 27, 2015

To: the Selection Committee for the Alfred Bader Award in Bioinorganic or Bioorganic Chemistry

It is a pleasure to nominated R. David Britt for this award. In his 27 years as a Professor of Chemistry at UC Davis, R. David Britt has run a strong and well-funded program applying electron paramagnetic resonance methods to study the roles of metals and radicals in metalloenzymes and bioinspired catalysts, with a particular interest in reactions involving conversion of solar energy to chemical energy. In his Ph.D. work at UC Berkeley, under the tutelage of Dr. Melvin P. Klein in the Laboratory of Chemical Biodynamics, David had built a pulse EPR spectrometer and used it in pioneering electron spin echo envelope modulation (ESEEM) studies of the manganese cluster centered Oxygen Evolving Complex (OEC) of Photosystem II, for example characterizing the binding of the ammonia water oxidation inhibitor to the manganese cluster [20]. As a new Assistant Professor at UC Davis, David and his new students built a more capable second generation pulse EPR spectrometer [29], with a greater frequency range and pulse electron nuclear double resonance (ENDOR) capabilities. With this instrument they carried out a number of seminal experiments on the OEC, including demonstrating there are four exchange couple manganese ions arranged in a 3 + 1 "dangler Mn" geometry [77], characterizing the interaction between the Mn cluster and the PSII tyrosine radical YZ. [40,59], and determining the amino acid ligation of the cluster with its single key histidine ligand [35]. Professor Britt has maintained continual funding by NIH and DOE for this OEC EPR work for over 25 years.

Over this time, Professor Britt has developed a strong instrumentation suite in his CalEPR center, with both homebuilt and commercial EPR instruments covering a frequency range from 8 to 130 GHz, along with a unique high power 263 GHz spectrometer currently under construction. With these instruments the Britt lab has expanded their focus beyond PSII, working on a wide variety of enzymes and catalytic synthetic complex. For example, building on research experience in PSII water splitting, the lab has tackled EPR characterization of Nocera's electrodeposited cobalt water oxidation catalyst and multinuclear cobalt cluster analogs [137, 151] along with a wider array of catalysts developed in the NSF funded Solar Fuels CCI (Gray, PI).

The Britt lab has also used these multifrequency advanced EPR instruments to characterize a wide variety of metal and radical based enzyme mechanisms [124, 139, 152, 158, 188], including catalytic RNA "ribozyme" systems [110]. One particular interesting problem recently tackled by the Britt group is the bioassembly of the catalytic "H-Cluster" of the [Fe-Fe] hydrogenase family. This six-Fe cluster, which can form ~10,000 H₂ molecules/sec, consists of a [4Fe-4S] cluster bridged to a unique dinuclear

Fe center which has CO and CN ligands along with an azadithiolate bridge between the two irons. The CO and CN are sourced from tyrosine, and the Britt lab and their collaborators recently showed in a series of papers that instead of the CO and CN ligands binding to a preformed 2Fe-2S cluster, instead a radical generating enzyme HydG cleaves the tyrosine substrate and binds the resultant diatomics on a unique fifth Fe of a novel five Fe cluster [173, 174, 192]. Intermediates of this enzyme cycle have been characterized for the first time with a variety of spectroscopic techniques, including EPR, stopped flow FTIR, and Mossbauer spectroscopy. Professor Britt and colleagues propose that the product of the HydG reaction is an Fe(CO)₂(CN)(l-cysteine) organometallic synthon [192]. In their model two of these synthons are converted into the binuclear cluster via their interactions with two other maturase enzymes, HydF and HydE, through processes currently being investigated by the Britt group and their collaborators.

This is novel chemistry and is gaining significant attention in the metallobiochemistry community. Because of his interesting scientific advancements, Professor Britt receives numerous lecture invitations each year. He is known for giving clear and enthusiastic lectures about his research. Along these lines I note that in 2014 Professor Britt received the International EPR Society Gold Medal Award, which is awarded once every 3-years, and he is the recipient of the 2016 Royal Society of Chemistry Bruker Prize for career contributions to EPR spectroscopy. He was also recently elected Chair of the Metals in Biology Gordon Research Conference (2017).

Sincerely,

Alan L. Balch

Distinguished Professor

Alan L Bell

R. David Britt

Distinguished Professor of Chemistry University of California, Davis

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Education:

B.S. Physics, North Carolina State University, Raleigh NC 1978

Ph.D Physics, University of California, Berkeley, CA 1988

Professional Positions:

1989-1994	Assistant Professor, Department of Chemistry, University of California at Davis
1994-1997	Associate Professor, Department of Chemistry, University of California at Davis
1997-	Professor, Department of Chemistry, University of California at Davis
2005-2008	Chair, Department of Chemistry, University of California at Davis

Honors and Awards:

1978-81	National Science Foundation Graduate Fellowship, University of California, Berkeley
1989	Camille and Henry Dreyfus New Faculty Award
1995	International EPR Society Young Investigator Award
2012	Elected Fellow of the American Association for the Advancement of Science
2012	Japan Society for the Promotion of Science Visiting Fellow
2014	International EPR Society Gold Medal (awarded every third year)
2015	University of California Distinguished Professor
2016	Royal Society of Chemistry Bruker Prize

Professional Service:

2000-04	Member, National Institutes of Health Metallobiochemistry Study Section
2006	Chair, Photosynthesis Gordon Research Conference
2008-13	Scientific Advisory Board, NSF Center for Chemical Innovation in Solar Fuels.
2010-13	Elected Member, National High Magnetic Field Laboratory EMR Advisory Committee
2012-	External Advisory Committee, ASU Center for Bio-Inspired Solar Fuel Production
2014	Elected Chair of the 2017 Metals in Biology Gordon Research Conference
2014-	External Advisory Committee, National High Magnetic Field Laboratory

LIST OF PUBLICATIONS

R. David Britt

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 Waves 2:1083-1096.
- Goodin, D. B., V. K. Yachandra, R. D. Britt, K. Sauer, and M. P. Klein. Light-Induced Changes in X-ray Absorption (K-edge)
 Energies of Manganese in Photosynthetic Membranes.
 Biochim. Biophys. Acta 767:209-216.
- 3. 1984 Goodin, D. B., V. K. Yachandra, R. D. Guiles, R. D. Britt, A. E. McDermott, K. Sauer, and M. P. Klein. Light-Induced Changes in X-ray Absorption (K-edge) Energies of Manganese in Photosynthetic Membranes. In EXAFS and Near Edge Structure III (eds. K. O. Hodgson, B. Hedman, and J. E. Penner-Hahn) Springer-Verlag, New York. p.130-135.
- 4. 1986 Yachandra, V. K., R. D. Guiles, A. E. McDermott, R. D. Britt, K. Sauer, and M. P. Klein. Structure of the Manganese Complex in Photosystem II Studied Using EXAFS Spectroscopy. The S₁ State of the O₂-Evolving Complex in Spinach. Biochim. Biophys. Acta 850:324-332.
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- 6. 1987 Britt, R. D. and M. P. Klein. A Versatile Loop-Gap Resonator Probe for Low-Temperature Electron Spin-Echo Studies. J. Magn. Reson. 74:535-540.
- 7. 1987 <u>Britt, R. D.</u>, K. Sauer and M. P. Klein. **Electron Spin Echo Studies of PSII Membranes.** *In Progress in Photosynthesis Research* (ed. J. Biggins) Martinus Nijhoff, Dordrecht. Vol <u>1</u>:573-576.
- 8. 1987 Yachandra, V. K., R. D. Guiles, A. E. McDermott, J. Cole, <u>R. D. Britt</u>, S. L. Dexheimer, K. Sauer, and M. P. Klein. **Comparison of**

- the Structure of the Mn-Complex in the S₁ and S₂ States of the Photosynthetic O₂-Evolving Complex. An X-ray Absorption Spectroscopy Study. Biochemistry <u>26</u>:5974-5981.
- 9. 1987 Cole, J., V. K. Yachandra, R. D. Guiles, A. E. McDermott, R. D. Britt, S. L. Dexheimer, K. Sauer, and M. P. Klein. Assignment of the g=4.1 EPR Signal to Mn in the S₂ State of the Photosynthetic O₂-Evolving Complex: An X-ray Absorption Edge Spectroscopy Study. Biochim. Biophys. Acta 890:395-398.
- 10. 1987 Cole, J., V. K. Yachandra, A. E. McDermott, R. D. Guiles, R. D. Britt, S. L. Dexheimer, K. Sauer, and M. P. Klein. Structure of the Mn Complex of Photosystem II upon Removal of the 33 kDa Extrinsic Protein: An X-ray Absorption Spectroscopy Study. Biochemistry 26:5967-5973.
- 11. 1987 McDermott, A. E., R. D. Guiles, V. K. Yachandra, J. Cole, R. D. Britt, S. L. Dexheimer, K. Sauer, and M. P. Klein. **X-ray Absorption Spectroscopy of Manganese and Iron in the Photosynthetic Apparatus.** In *Biophysics and Synchrotron Radiation* (ed. A. Bianconi, and A. G. Castellano) Springer-Verlag, Heildelberg. p. 223-230.
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- 14. 1987 McDermott, A. E., V.K. Yachandra, R. D. Guiles, R. D. Britt, S. L. Dexheimer, K. Sauer, and M. P. Klein. Characterization of the Mn-Containing O₂-Evolving Complex from the Cyanabacterium Synechococcus Using EPR and X-ray Absorption Spectroscopy. In Progress in Photosynthesis Research (ed. J. Biggins) Martinus Nijhoff, Dordrecht. Vol 1:565-568.
- 15. 1987 McDermott, A. E., V. K. Yachandra, R. D. Guiles, <u>R. D. Britt</u>, S. L. Dexheimer, K. Sauer, and M. P. Klein. **Iron X-ray Absorption**

- **Spectroscopy of Low Potential Acceptors in Photosystem I.** In *Progress in Photosynthesis Research* (ed. J. Biggins) Martinus Nijhoff, Dordrecht. Vol 1:249-252.
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- 18. 1988 Britt, R. D., J. L. Zimmermann, K. Sauer, and M. P. Klein. **ESEEM**Studies of Ammonia Binding to the Catalytic Mn of the Water

 Oxidation Complex of Photosystem II. In Pulsed EPR: A New

 Field of Applications. (eds. C. P. Keijzers, E. J. Reijerse, and J. Schmidt) Nort Holland, Amsterdam. pp. 150-155.
- 19. 1988 Sauer, K., R. D. Guiles, A. E. McDermott, J. L. Cole, V. K. Yachandra, J. L. Zimmermann, M. P. Klein, S. L. Dexheimer, and R. D. Britt. Spectroscopic Studies of Mn Involvement in Photosynthetic O₂ Evolution. Chemica Scripta 28A:87-91.
- 20. 1989 Britt, R. D., J. L. Zimmermann, K. Sauer, and M. P. Klein.

 Ammonia Binds to the Catalytic Mn of the Oxygen Evolving

 Complex of Photosystem II: Evidence by Electron Spin Echo

 Envelope Modulation Spectroscopy. J. Am. Chem. Soc.

 111:3522-3532.
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- and S₁ States Determined by X-ray Absorption Spectroscopy. Biochemistry 29:486-496.
- 23. 1990 Britt, R. D., V. J. DeRose, V. K. Yachandra, D. K. Kim, K. Sauer, and M. P. Klein. Pulsed EPR Studies of the Manganese Center of the Oxygen-Evolving Complex of Photosystem II. In Current Research in Photosynthesis, Vol. 1, ed., M. Baltscheffsky, Kluwer Academic, pp. 769-772.
- 24. 1990 Kim, D. H., R. D. Britt, M. P. Klein, and K. Sauer. The g=4.1 EPR Signal of the S₂ State of the Photosynthetic Oxygen Evolving Complex Arises from a Multinuclear Mn Cluster. J. Am. Chem. Soc. 112:9389-9391.
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- 29. 1992 Sturgeon, B. E. and R. D. Britt. **Design of a Sensitive Pulsed EPR Spectrometer with an 8 to 18 GHz Frequency Range.** Rev. Sci. Instrum. 63:2187-2192.

- 30. 1992 Britt, R. D. and M. P. Klein. Electron Spin Echo Envelope Modulation Studies of Mixed Valence Manganese Complexes:
 Applications to the Catalytic Manganese Cluster of Photosynthetic Oxygen Evolution. In Pulsed Magnetic Resonance: NMR, ESR, and Optics, a recognition of E.L. Hahn (ed. D. M. S. Bagguley) Clarendon Press, Oxford pp. 390-410.
- 31. 1992 <u>Britt, R. D.</u>, G. A. Lorigan, K. Sauer, M. P. Klein, and J. L. Zimmermann. The g=2 multiline EPR signal of the S₂ state of the photosynthetic oxygen-evolving complex originates from a ground spin state. Biochim. Biophys. Acta 1040:95-101.
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 Quinone Electron Acceptor, Q_A, of PSII. In Photosynthesis: from Light to Biosphere (ed. P. Mathis) Kluwer Academic Publishers, Amsterdam, Vol. I, pp. 775-778.
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- 44. 1995 Diner, B. A., M. Zheng, X. S. Tang, G. C. Dismukes, D. A. Force, D. W. Randall, and R. D. Britt. Environment and Function of the Redox Active Tyrosines of Photosystem II. In Photosynthesis: from Light to Biosphere (ed. P. Mathis) Kluwer Academic Publishers, Amsterdam, Vol. II, pp. 229-234.
- 45. 1995 Force, D. A., D. W. Randall, R. D. Britt, X.-S. Tang, and B. A. Diner. ²H ESE-ENDOR Study of Hydrogen Bonding to the Tyrosine Radicals Y_D and Y_Z of Photosystem II. J. Am. Chem. Soc. 117:12643-12644.
- 46. 1996 Britt, R. D. Electron Spin Echo Methods in Photosynthesis Research. In Advances in Photosynthesis: Biophysical Techniques in Photosynthesis (eds. A.J. Hoff and J. Amesz) Kluwer Academic Publishers, Amsterdam, pp. 235-253.

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- 50. 1997 Campbell, K. A., J. M. Peloquin, B. A. Diner, X.-S. Tang, D. A. Chisholm, and R. D. Britt. The τ-Nitrogen of D2 Histidine 189 is the Hydrogen Bond Donor to the Tyrosine Radical Y_D of Photosystem II. J. Am. Chem. Soc. 119:4787-4788.
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- 159. 2012 Stoll, S., Y.-T. Lee, M. Zhang, R. F. Wilson, R. D. Britt, and D. B. Goodin. Double Electron-Electron Resonance Shows

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- 160. 2012 Harley, S. J., H. E. Mason, J. G. McAlpin, R. D. Britt, and W. H. Casey. A ³¹P NMR Investigation of the CoPi Water-Oxidation Catalyst. Chem. Eur. J. 18:10476-10479.
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 Cofactor of Class Ib Ribonucleotide Reductase: Enzymatic
 Generation of Superoxide Is Required for Tyrosine Oxidation
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- 171. 2013 Stich, T. A., J. Krzystek, B. Q. Mercado, J. G. McAlpin, C. A. Ohlin, M. M. Olmstead, W. H. Casey, and R. D. Britt. Structural Insights Into [Co₄O₄(C₅H₅N)₄(CH₃CO₂)₄]⁺, A Rare Co(IV)-Containing Cuboidal Complex. Polyhedron, 64:304-307.
- 172. 2013 Najafpour, M.M., M. Kompany-Zareh, A. Zahraei, D. J. Sedigh, H. Jaccard, M. Khoshkam, R. D. Britt and W. H. Casey. Mechanism, Decomposition Pathway And New Evidence For Self-Healing Of Manganese Oxides As Efficient Water Oxidizing Catalysts: New Insights. Dalton Trans. 42:14603-14611.
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- 176. 2014 Britt, R. D. and Paul H. Oyala. One Step Closer to O₂. Science, 345:736.
- 177. 2014 Stich, T. A., W. K. Myers, and R. D. Britt. Paramagnetic Intermediates Generated by Radical S-Adenosylmethionine (SAM) Enzymes. Acc. Chem. Res. 47:2235-2243.
- 178. 2014 Myers, W. K., T. A. Stich, D. L. M. Suess, J. M. Kuchenreuther, J. R. Swartz, and R. D. Britt. The Cyanide Ligands of [FeFe] Hydrogenase: Pulse EPR Studies of ¹³C and ¹⁵N-Labeled H-Cluster. J. Am. Chem. Soc. 136:12237-12240.
- 179. 2014 Karagas, N. E. C. N. Osborn, D. J. Osborn, A. L. Dzierlenga, P. Oyala, M. E. Konkle, E. M. Whitney, R. D. Britt, and L. M. Hunsicker-Wang. The Reduction Rates Of DEPC-Modified Mutant Thermus Thermophilus Rieske Proteins Differ When There Is a Negative Charge Proximal to the Cluster. J. Biol. Inorg. Chem. 19:1121-1135.
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- Stich, T. A. and <u>R. D. Britt</u>. **Advanced Electron Paramagnetic Resonance Studies of the Oxygen Evolving Complex.** In *Metalloproteins: Theory, Calculations, and Experiments.* A. E. Cho and W. Goddard, Eds., CRC Press. pp. 1-58.
- Oyala, P. H., T. A. Stich, and R. D. Britt. Metal Ion Oxidation State Assignment Based on Coordinating Ligand Hyperfine Interaction. Photosynth. Res., 124:7-18.
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- 184. 2015 Gagnon, D. .M, M. B. Brophy, S. E. J. Bowman, T. A. Stich, C. L. Drennan, R. D. Britt, and E. M. Nolan. Manganese Binding Properties of Human Calprotectin Under Conditions of High and Low Calcium. J. Am. Chem. Soc. 137:3004-3016.

- Parthasarathy, A., T. A. Stich, S. T. Lohner, A. Lesnefsky, M. Fincker, R. D. Britt, and A. M. Spormann. Biochemical and EPR-spectroscopic Investigation into Heterologously Expressed Vinyl Chloride Reductive Dehalogenase (VcrA) from Dehalococcoides mccartyi strain VS. J. Am. Chem. Soc. 137:3525-3532.
- 186. 2015 Britt, R. D., D. L. M. Suess, and T. A. Stich. An Mn(V)-Oxo Role in Splitting Water? Proc. Natl. Acad. Sci. U.S.A. 112:5265-5266.
- Oyala, P. H., T. A. Stich, R. J. Debus, and R. D. Britt. Ammonia Binds to the Dangler Manganese of the Photosystem II Oxygen-Evolving Complex. J. Am. Chem. Soc. 137:8829-8837.
- 188. 2015 Chang, Y. G., S. Cohen, C. Phong, W. K. Meyers, Y. I. Kim, R. Tseng, J. Lin, L. Zhang, J. Boyd, R. D. Britt, M. Rust, S. S. Golden, and A. LiWang. **KaiB Fold Switching Is The Linchpin That**Joins Oscillator Function To Clock Output. Science 349:324-328.
- 189. 2015 Suess, D. L. M. and R. D. Britt. **EPR Spectroscopic Studies of** [FeFe]-Hydrogenase Maturation. Top. Catal. 58:699-707.
- 190. 2015 Tao, L., T. A. Stich, C. N. Butterfield, C. A. Romano, T. G. Spiro, B. M. Tebo, W. H. Casey, and R. D. Britt. Mn(II) Binding and Subsequent Oxidation by the Multicopper Oxidase MnxG Investigated by Electron Paramagnetic Resonance Spectroscopy. J. Am. Chem. Soc. 137:10563-10575.
- 191. 2015 Butterfield, C. N, L. Tao, K. N. Chacon, T. G. Spiro, N. J. Blackburn, W. H. Casey, R. D. Britt, B. M. Tebo. Multicopper Manganese Oxidase Accessory Proteins Bind Cu and Heme. Biochim. Biophys. Acta, published online before print: doi:10.1016/j.bbapap.2015.08.012
- 192. 2015 Suess, D. L. M., I Burstel, L. De La Paz, J. M. Kuchenreuther, C. Pham, S. P. Cramer, J. R. Swartz, and R. D. Britt. Cysteine as a Ligand Platform in the Biosynthesis of the FeFe Hydrogenase H-cluster. Proc. Natl. Acad. Sci. U.S.A., Published online before print, doi: 10.1073/pnas.1508440112

R. David Britt: Grants (1997-present [since promotion to Professor])

Research Grants

1. Title: Time-Domain EPR Studies of Tyrosine Z of Photosystem II

Agency: National Science Foundation

Amount: \$243,806 Direct Cost; \$290,000 Total Cost

Dates: 2/1/96 - 1/31/99 PI: R. David Britt

2. Title: Pulsed EPR Studies of Biological Manganese Clusters

(R01 GM48242)

Agency: National Institutes of Health

Amount: \$507,547 Direct Cost; \$706,233 Total Cost

Dates: 7/1/97 - 6/30/01 PI: R. David Britt

3. Title: Pulsed EPR Studies of Photosystem II Radicals

Agency: National Science Foundation

Amount: \$280,148 Direct Cost; \$375,000 Total Cost

Dates: 3/15/99 - 2/28/02 PI: R. David Britt

4. Title: EPR Studies of Cytochrome c Oxidase Intermediates

Agency: National Institutes of Health (R21)

Amount: \$140,000 Direct Cost: \$196,666 Total Cost

Dates: 4/1/00 - 3/31/02 PI: R. David Britt

5. Title: EPR Studies of Metal-Nucleic Acid Interactions

Agency: National Institutes of Health (R01 GM61211) Amount: \$687,092 Direct Cost; \$915,459 Total Cost

Dates: 4/1/00 - 3/31/04 PI: R. David Britt

6. Title: Pulsed EPR Studies of Biological Manganese Clusters

Agency: National Institutes of Health (R01 GM48242) Amount: \$720,000 Direct Cost; \$980,791 Total Cost

Dates: 7/1/01 - 6/30/06 PI: R. David Britt

7. Title: Multifrequency EPR/ENDOR Spectrometer

Agency: National Institutes of Health (S01 RR021075) Amount: \$500,000 Direct Cost; \$500,000 Total Cost

Dates: 1/15/05 – 1/14/07 PI: R. David Britt Co-PIs: Glenn Millhauser, John Voss, Steve Cramer, Michael Marletta, Judith

Klinman, Paul Ludden, Edward Solomon

8. Title: EPR Studies of Biological PCET Elements

Agency: National Institutes of Health (R01 GM073789) Amount: \$706,601 Direct Cost; \$1,042,207 Total Cost

Dates: 3/5/05 - 2/28/11 PI: R. David Britt

9. Title: Pulsed EPR Studies of Biological Manganese Clusters

Agency: National Institutes of Health (R01 GM48242) Amount: \$850,227 Direct Cost; \$1,187,131 Total Cost

Dates: 7/1/06 - 5/31/12 PI: R. David Britt

10. Title: A High Magnetic Field Mössbauer Instrument

Agency: National Institutes of Health (1S10 RR023656)

Amount: \$273,000 Direct & Total Cost

Dates: 4/15/08 - 4/14/09 PI: Stephen P. Cramer Co-PI: R. David Britt et al.

11. Title: EPR Spectroscopy of the Cobalt Water Splitting Catalyst

Agency: National Science Foundation EAGER (CHE-0939178)

Amount: \$165,226 Direct Cost; \$240,000 Total Cost

Dates: 8/1/09 - 7/31/12 PI: R. David Britt

12. Title: Purchase of an X-ray Diffractometer

Agency: National Science Foundation (08-002425) ARRA

Amount: \$348,552 Direct Cost; \$349,828 Total Cost

Dates: 08/01/2009 - 07/31/2012

PI: R. David Britt (as Chemistry Dept Chair)

13. Title: Pulsed EPR Studies of Biological Manganese Clusters

Agency: National Institutes of Health (R01, GM48242, ARRA Supplement)

Amount: \$100,000 Direct Cost; \$153,000 Total Cost

Dates: 7/1/10 - 6/30/11 PI: R. David Britt

14. Title: Multifrequency Pulsed EPR Studies of the Photosystem II Oxygen

Evolving Complex

Agency: Department of Energy, Basic Energy Sciences Amount: \$360,000 Direct Cost; \$540,000 Total Cost

Dates: 9/1/11 – 8/30/14 PI: R. David Britt 15. Title: Photoreceptor Regulation and Optimization of Energy Harvesting in

Nostoc punctiforme

Agency: Department of Energy SISGR (DESC0002395)

Amount: \$261,327 Direct Cost; \$387,970 Total Cost (Britt portion)

Dates: 9/15/09 - 9/14/12 PI: J. Clark Lagarias Co-PI: R. David Britt et al.

16. Title: Membrane Organized Chemical Photo Redox Systems

Agency: Department of Energy (DESC0004334)
Amount: \$65,271 Direct Cost; \$100,000 Total Cost

Dates: 5/1/10 - 4/30/13 PI: R. David Britt

17. Title: Acquisition of Dual Frequency Electron Paramagnetic Resonance

Spectrometer

Agency: National Science Foundation (CRIF) CHE-1048671

Amount: \$301,168 Direct & Total Cost

Dates: 12/15/10 - 11/30/13

PI: Carlito B. Lebrilla (as Chemistry Dept Chair)

Co-PI: R. David Britt

18. Title: Mechanisms of Co, Ni, and Mn based Water Splitting Catalysts

Probed by Advanced EPR Spectroscopy

Agency: National Science Foundation (CHE-1213699) Amount: \$341,233 Direct Cost; \$449,999 Total Cost

Dates: 6/1/12 - 5/31/15 PI: R. David Britt

19. Title: Mechanistic Studies of the Radical SAM Enzyme QueE

Agency: NIH – National Institute of General Medical Sciences

(supplement to R01 GM072623)

Amount: \$85,454 Direct Cost; \$131,599 Total Cost (Britt portion)

Dates: 5/1/12 - 4/30/15 PI: Vahe Bandarian Co-PI: R. David Britt

20. Title: Photoreceptor Regulation and Optimization of Energy Harvesting in

Nostoc punctiforme

Agency: Department of Energy SISGR (DESC0002395)

Amount: \$71,808 Direct Cost; \$110,584 Total Cost (Britt portion)

Dates: 9/15/12 - 9/14/13
PI: J. Clark Lagarias
Co-PI: R. David Britt et al.

21. Title: Chemical Bonding Center: Chemistry as the Driver for

Transformative Research and Innovation

Agency: NSF Center for Chemical Innovation Solar Fuels

Amount: \$524,272 Direct Cost; \$810,000 Total Cost

Dates: 10/1/13 – 9/30/18 PI: Harry Gray (Cal Tech)

Co-PI: R. David Britt

22. Title: Mechanisms of Radical SAM Enzymes Probed by EPR Spectroscopy

Agency: National Institutes of Health (R01 GM104543) Amount: \$900,000 Direct Cost; \$1,399,500 Total Cost

Dates: 7/1/14 - 6/30/18 PI: R. David Britt

23. Title: MRI: 260 GHz Pulse EPR/DEER Spectrometer

Agency: National Science Foundation

Amount: \$1,435,026 Direct Cost; \$678,997 Cost Share; \$2,263,323 Total Cost

Dates: 7/1/14 - 6/30/17 PI: R. David Britt

Co-PI: Neville Luhmann, Jr.

24. Title: Multifrequency Pulsed EPR Studies of the Photosystem II Oxygen

Evolving Complex

Agency: Department of Energy, Basic Energy Sciences Amount: \$419,344 Direct Cost; \$626,576 Total Cost

Dates: 7/1/15 - 6/30/17PI: R. David Britt

Seminars and Chaired Sessions (since advancement to Professor Step VI in 2007):

NSF Chemical Bonding Center "Powering the Planet" Workshop (2007) Ventura, CA

Boise State University, Chemistry Department Seminar (2008) Boise ID

Protein Cofactors, Radicals and Quinones Gordon Research Conference (2008) Ventura CA

Iron Sulfur Enzymes Gordon Research Conference (2008) Colby-Sawyer College, New London NH

International Conference on Magnetic Resonance in Biological Systems (2008) San Diego, CA

Photosynthesis Gordon Research Conference (2009) Bryant University, Smithfield, RI

International Meeting of the Royal Society of Chemistry EPR group (2009) Norwich, UK

Session Chair: International Meeting of the Royal Society of Chemistry EPR group (2009) Norwich, UK

North American/Greece/Cyprus Workshop on Paramagnetic Materials (2009) Cyprus

NSF Center for Chemical Innovation Solar Fuels Annual Retreat [workshop participant] (2009) Ventura, CA

ACERT Workshop on Structure and Dynamics by Multifrequency ESR/EPR (2009) Cornell University, Ithaca NY

Session Chair: Second Annual Workshop of the 10+10 Alliance (2009) Davis CA,

Session Chair: International Conference on Tetrapyrrole Photoreceptors of Photosynthetic Organisms (2009) Asilomar CA

Institute for Complex Adaptive Matter Annual Meeting (2010) Davis CA

NSF Center for Chemical Innovation Solar Fuels Annual Retreat (2010) Huntington Beach CA

University of Oregon, Chemistry Department Seminar (2010) Eugene OR

Chemistry and Biology of Tetrapyrroles Gordon Research Conference (2010) Newport RI

Session Chair: Western Regional Photosynthesis Conference (2010) Asilomar CA

Session Chair (X2): International Conference on Magnetic Resonance in Biological Systems (2010) Cairns Australia

Discussion Leader: Research Frontiers in Bioinspired Energy Workshop, National Academy of Sciences (2010) Washington DC

International Conference on Magnetic Resonance in Biological Systems (2010) Cairns Australia

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Metals in Biology Gordon Research Conference (2011) Ventura CA

American Chemical Society National Meeting, Symposium on "Magnetic Spectroscopic Approaches to the Study of Metals in Biology" (2011) Anaheim CA

Solar Fuels: Light Capture and Electron Flow, J. Heyrovsky Institute of Physical Chemistry (2011) Prague, Czech Republic

Brian Hoffman 70th Birthday Symposium (2011) Vancouver, British Columbia

Session Chair: International Congress on Bioinorganic Chemistry (ICBIC-15) (2011) Vancouver, British Columbia

40th Southeastern Magnetic Resonance Conference, Plenary Lecture (2011) Atlanta, GA,

Inorganic Chemistry special student lecture (invited by Prof. Harry Gray) California Institute of Technology (2012) Pasadena CA

American Chemical Society National Meeting, Symposium on "Photochemistry in Biology" (2012) Philadelphia, PA

NSF CCI Solar Site Visit California Institute of Technology (2012) Pasadena CA. Chaired reviewers' question and answer session.

DOE Physical Biosciences Research Meeting (2012) Baltimore MD

Chemistry Department Seminar, Marquette University (2012) Milwaukee WI

Yamaguchi University (2012) Yamaguchi Japan. Japan Society for the Promotion of Science Lecture

Japan Society of Electron Spin Science and Technology (2012) Sapporo Japan. Plenary Lecture

Hokkaido University (2012) Sapporo Japan. Japan Society for the Promotion of Science Lecture

Tohoku University (2012) Sendai Japan. Japan Society for the Promotion of Science Lecture

Nagoya University (2012) Nagoya Japan. Japan Society for the Promotion of Science Lecture

Kobe University (2012) Kobe Japan. Japan Society for the Promotion of Science Lecture

Bioinorganic Chemistry Gordon Research Seminar (2013) Ventura CA. Discussion Leader

MIT/Harvard Joint Inorganic Seminar (2013) Cambridge MA

International Society of Magnetic Resonance International Conference (2013) Rio de Janeiro Brazil

Rocky Mountain Conference on Magnetic Resonance (2013) Denver CO

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Solar Fuels Gordon Research Conference (2014) Ventura CA. Discussion Leader

Dean Winston Ko Retirement Symposium (2013) Davis CA

Texas A& M University Chemistry Seminar (2013) College Station TX

University of Michigan Chemistry Seminar (2013) Ann Arbor MI

National Renewable Energy Laboratory Seminar (2013) Golden CO

CBIG Chemical Biology Seminar (2013) Davis CA

DOE Photosynthesis Research Meeting (2012) Annapolis MD

University of Florida Chemistry Seminar (2013) Gainesville FL

Florida State University Chemistry Seminar (2013) Tallahassee FL

Science in the Country (2013) Half Moon Bay CA

University of California, Berkeley Chemistry Seminar (2013) Berkeley CA

California Institute of Technology Chemistry Seminar (2014) Pasadena CA

Inorganic Chemistry special student lecture (invited by Prof. Harry Gray) California Institute of Technology (2014) Pasadena CA

Solar Fuels Gordon Research Conference (2014) Ventura CA. Discussion Leader

Metals in Biology Gordon Research Conference (2014) Ventura CA.

Stanford University Chemistry Seminar (2014) Palo Alto CA

Emory University Chemistry Seminar (2014) Atlanta GA

Georgia State University Chemistry Seminar (2014) Atlanta GA

47th Annual International Meeting of the ESR Spectroscopy Group of the Royal Society of Chemistry Keynote Lecture (2014) Dundee Scotland UK

University of Washington Chemistry Seminar (2014) Seattle WA

97th Canadian Chemistry Conference (2014) Vancouver Canada

Frontiers in Metallobiochemistry III (2014) Penn State University, PA

Iron-Sulfur Enzymes Gordon Research Conference (2014) Easton MA

Photosynthesis Gordon Research Conference Keynote Lecture (2014) West Dover VT

Catalytic Systems for Chemical Energy Conversion International Symposium (2014) Mulheim Germany

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European Federation of EPR Groups Plenary Lecture (2014) Marseilles France

Portland State University - Oregon Health Sciences University Joint Seminar (2014)

Joint Conference of Asia-Pacific EPR/ESR Symposium/ International EPR Society Gold Medal Award Lecture (2014) Nara Japan

University of Pennsylvania Biological Chemistry Seminar (2014) Philadelphia PA

University of Illinois Biochemistry Seminar (2014) Urbana-Champaign IL

Oxford University Departmental Seminar (2014) Oxford UK

Imperial University Special Seminar (2014) London UK

Queen Mary University Departmental Seminar (2014) London UK

University of California, Irvine Inorganic Chemistry Seminar (2015) Irvine CA

University of Florida, Departmental Seminar (2015) Gainesville FL

National High Magnetic Field Laboratory, Future Lab Directions Workshop (2015) Tallahassee FL

University of California Davis, Biophysics Seminar (2015) Davis CA

Resolv EPR Symposium (2015) Bochum Germany

Sacramento State University Chemistry Seminar (2015) Sacramento CA

Williamette University Chemistry Seminar (2015) Salem OR

Scheduled Invited Seminars and Chaired Sessions:

Southeastern Magnetic Resonance Conference (2015) Daytona Beach FL

University of California Santa Barbara, Departmental Seminar (2015) Santa Barbara CA

University of California Riverside, Departmental Seminar (2015) Santa Barbara CA

University of Utah, Departmental Seminar (2015) Salt Lake City UT

University of California Merced, Departmental Seminar (2015) Merced CA

Frontiers of Iron Chemistry in Biology Symposium, Pacifichem (2015) Honolulu HI

Metals in Biology Gordon Research Conference (2016) Ventura CA

American Society of Biochemistry and Molecular Biology National Meeting (2016) San Diego CA

International Hydrogenase Conference (2016) Marseille France

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79th Harden Conference: Oxygen Evolution and Reduction (2016) Innsbruck, Austria

American Chemical Society National Meeting, Symposium on "Understanding Enzymatic Catalysis across Multiple Timescales: Experiment and Theory" (2016) San Diego CA

American Chemical Society National Meeting, (2016) Philadelphia PA

Royal Society of Chemistry Bruker Prize Lecture, (2016) Colchester UK

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R. David Britt: 20 SIGNIFICANT PUBLICATIONS

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Professor **David Britt**, Ltr of Support, ACS Alfred Bader Award in Bioinorganic or Bioorganic Chemistry

A flagship of Professor Britt's program is the implementation of a bewildering array of EPR techniques to unveil the structure of biological cofactors. A decade before the Photosystem II (PSII) X-ray crystal structures by Barber (2004), Professor Britt applied electron spin echo electron-nuclear double resonance to show that the tyrosine EPR signal of PSII was altered relative to that observed in a manganese-depleted preparation owing to a magnetic interaction between the manganese cluster of PSII and a nearby tyrosine radical. Not only did he show they were proximate but he was even able to accurately decipher the distance between the two cofactors. It is now known that the tyrosine radical is the key residue that participates in a proton-coupled electron transfer reaction to relay hole equivalents into the OEC for water oxidation. That Britt was able to make this prediction in the mid-1990s when there were only rudimentary 2D electron diffraction maps of the photosynthetic membrane (where the OEC was invisible) is truly stunning. And if that were not enough, he then went on to provide one of the great biophysics studies in my experience. The structure of the OEC was a point of controversy dating back to the 1960s. Britt went against the dogma in the field and boldly proposed a 3+1 Mn structure with the lone manganese labeled by him as "the dangler". His model was later confirmed the membrane-bound X-ray structures. I am still amazed that Britt was able to deduce the structure on the basis of EPR, long before the structural biologists arrived on the scene. His prediction was bold and striking, and better yet, he was entirely correct; I view this work as a gold standard in science.

In addition to PSII, Britt has impacted other areas of bioinorganic chemistry. I briefly mention the SAM enzymes where he has identified almost every possible intermediate, from an analogue of the initial 5'-dA radical to the product radical L-beta-lysine. Characteristic of the Britt signature, the work is meticulously performed and overpowering in its detail and insight.

David Britt is THE premier scientist in the world today in EPR spectroscopy and he has provided the scientific community with the some of the most penetrating and insightful biophysical studies of this generation. I enthusiastically support the nomination of Professor Britt for the ACS Bader Award.

With best regards,

Daniel G. Nocera

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Britt has made seminal contributions through his development and use of advanced pulsed EPR, (ESEEM, ENDOR) methods for our understanding of complex biological enzymatic mechanisms including: the O₂ evolving complex, the maturation of the cofactor essential for the FeFe hydrogenase (HydA), and many enzymes that utilize S-adenosylmethionine (SAM) and [4Fe4S]¹⁺ to initiate unprecedented radical mediated transformations. Bioinformatics studies have identified >100,000 enzymes that use SAM and [4Fe4S]¹, now classified as Radical SAM Superfamily (SF) members. Study of the detailed mechanisms of the transformations by these enzymes requires sophisticated protein purification methods, use of kinetics, isotopic labeling of the enzyme and substrate, and a tool box of time-resolved biophysical spectroscopies.

The Britt and Swartz labs have played a central role in our current understanding of the mechanism by which the unusual metallo-cofactor of the HydA, that catalyzes the interconversion of H₂ and H⁺ at 10⁴ s⁻¹, is biosynthesized. These enzymes contain a catalytic 6Fe-cluster, which is now known to be composed of a "regular" [4Fe4S] cluster bridged by a cysteine thiolate to a uniquely decorated 2Fe-subcluster. The two irons in this subcluster are bridged by dithiomethylamine and CO and each Fe is additionally coordinated to a CO and a CN-. The cofactor maturation process requires three proteins, HydE, HydG and HydF; the former two are members of the Radical SAM SF.

HydG was shown (Broderick, Peters and Roach) to use tyrosine as the substrate and produce p-creosol. Isotopic labeling studies of the tyrosine demonstrated that the atoms of its backbone are incorporated into all the CO and CN ligands. HydG also contains an auxiliary FeS cluster, as do many Radical SAM SF members. In a tour de force using protein biochemistry and spectroscopy (Science 2013 342, 472 and 2014, 343, 424), the Britt group in collaboration, identified the HydG bound organometallic Fe(CO)₂CN synthon and an organic radical intermediate that led to the favored mechanism for this amazing transformation.

While the details of how this synthon is transferred to HydA and acquires the dithiomethyamine ligand are still not understood, a recent paper (PNAS 2015 112, 11455) describes the spectroscopic evidence, that Cys likely binds to the auxiliary cluster and the organometallic synthon. A model for cluster assembly from this intermediate in the presence of HydE and HydF is proposed. The method of cell free protein expression, isotopic labeling and sophisticated spectroscopy is unraveling the amazing mechanisms by which complex and labile intermediates are made and used in Nature.