CURRICULUM VITAE

Edward Valerian Quadros, Ph.D.

Address:

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Biographical:

Date of Birth: November 27, 1946; India; U.S. Citizen

Educational Training:

1964 – 68, University of Poona, B.Sc. (Chemistry-Major)

1968 – 72, University of Bombay, M.Sc. (Applied Biology)

1973 – 76, University of London, Ph.D. (Biochemistry)

Positions:

2009-Present Research Professor, Departments of Medicine and Cell Biology

1996-2008 Associate Professor (Research), Departments of Medicine and Cell Biology,

SUNY-Downstate Medical Center, Brooklyn, NY 11203.

1984-1995 Assistant Professor (Research), Department of Medicine SUNY–Downstate Medical Center, Brooklyn, NY 11203

Positions Held:

1990 – 95, Research Biologist and Principal Investigator, Division of Research, V.A. Medical Center, Brooklyn, NY 11209

1984 – 90, Research Associate and Co-investigator on NIH project, Division of Hematology, V.A. Medical Center, Brooklyn, NY 11209

1979 – 83, Research Associate, Biochemistry Department, Scripps Clinic and Research Foundation, La Jolla, CA

- 1976 79, Research Associate, supported by the Welcome Trust, Department of Experimental Chemical Pathology, Westminster Hospital, London, SW1(UK).
- 1973 76, Research Fellow, Commonwealth Tropical Medicine Research Award, Department of Experimental Chemical Pathology, Westminster Hospital, London SW1 (UK).
- 1970 73, Research Assistant, project on Nutritional Disorders of the Nervous System for National Institutes of Health (U.S.A.), at Neuropathology Unit, J.J. Hospital and Grant Medical College, Bombay-8, India.
- 1968 69, Research Assistant, Glaxo project on Toxic Amblyopia at Neuropathology Unit, J.J. Hospital and Grant Medical College, Bombay-8, India.

Academic activities:

- 1. Served as Case based Learning (CBL) facilitator for GI and intermediary metabolism and for Cells to genes.
- 2. Lectured to graduate students on past and ongoing research in my laboratory and have been invited to speak at seminars organized by the Department of Anatomy and Cell biology, Department of Microbiology and Immunology and the Division of Hematology/Oncology. Currently there are two graduate students working in my laboratory towards a Ph.D. degree. In addition, a post doctoral fellow supported by the Swiss National Science Foundation spent one year on a joint project with Kent State University and my laboratory and a graduate student from the University of Reus, Spain, has spent 4 months in my laboratory to complete part of a collaborative research project.
- 3. Served as examiner for Ph.D proposals in the Departments of Biochemistry and Anatomy Cell biology.
- 4. Trained graduate students on laboratory rotation in various aspects of biochemical, cellular and molecular research techniques.
- 5. Accepted residents from the Department of Medicine and fellows from the Division of Hematology/Oncology and from Pediatric Hematology for training in basic science research.
- 6. In addition to the academic activities listed above, I have served as an external examiner of a Ph.D. thesis at the University of Nancy, France; University of Reus, Spain and have provided mentoring support to a post-doctoral fellow at the University of Utah, for collaborative work.

Current Teaching and Academic activities:

Medical School: Approximately > 100 hrs

- 1. First year Biochemistry lectures 3hours / second year lecture 1hr.
- 2. Case based learning:
 - a. Cells to genes, 18hrs

- b. GI and intermediary metabolism: 18hrs
- 3. Histology Laboratory: 65 hrs

Facilitators' discussions and preparations 2-4hrs

4. Interviewing applicants for medical school:

I interview one candidate every other week, typically 8-12 candidates per year.

Time allocation: 1hr for application review and 1-2hr for interview; total time, 16-24hrs

Graduate School: Approximately 80+ hrs

- 1. Course lecture: 6hrs
- 2. Thesis committees:

Thesis review at proposal, pre defense and final defense: Approximately 20hrs per candidate, 3 candidates per year, total 60hrs.

Thesis presentations and meetings, Approximately 6hrs per candidate, 3 candidates per year, total 18hrs

2008- Member, Executive Committee

2009-2010, Chairman, Research Resources Subcommittee

National and International Collaborations:

The research in my laboratory has involved collaboration with investigators at various institutions. These collaborations have included Axis-Shield, Norway; University of Nancy, France; University Aarhus, Denmark; Cleveland Clinic and Kent State University, Cleveland Ohio: University of Nebraska and Medarex Inc. In addition to continuing collaborations with these institutions I have established new collaborations with investigators at the University of Aachen, Germany; University of Reus, Spain; Trinity College, Ireland; The Human Genome Center, NIH; North Shore/Long Island Jewish Health System, NY and the Texas Genome Center, St Antonio.

National and International service:

As a laboratory recognized for its contribution to research in the vitamin B12 and folate fields, my laboratory serves as a worldwide reference center for biochemical and genetic disorders affecting pathways and genes involving these two nutrients. We receive samples for analysis from all over the world and are able to provide this service to patients affected by these rare disorders.

Commercialized Technology:

- 1. Holo transcobalamin assay: In an exclusive licensing agreement with Axis-Shield, a clinical diagnostics company based in the UK and Norway, I have been involved in the development of an assay for holo transcobalamin, a sensitive and accurate measure of vitamin B12 status. This assay is being marketed worldwide by Abbot Laboratories as part of the tests included in their new instrumentation platform. This test is expected to replace the current serum total B12 assay to diagnose B12 deficiency with a annual market of over 1 billion dollars. In addition to agreed royalty payments, the exclusive agreement includes the purchase and use of the monoclonal antibody to transcobalamin developed in my laboratory.
- 2. Folate receptor autoantibody assay: We have now signed an exclusive agreement with IMMCO Diagnostics, a New York based company specializing in immunodiagnostics to develop and market a assay for the identification of autoantibodies to folate receptors in serum. This autoantibody has a pathologic effect in a number of conditions such as neural tube defect pregnancy, infertility, and

miscarriages in women and cerebral folate deficiency in infancy. Intervention with pharmacological doses of folate can overcome many of the pathological effects of the autoantibody and therefore detection of this autoantibody and early intervention is critical to preventing permanent embryonic damage in pregnancy and neuro-developmental abnormalities in infants. The fairly high prevalence of this autoantibody in the elderly population could pose a risk factor for developing folate deficiency. The market potential for this test is predicted to be fairly large considering that every woman of child bearing age, every newborn and people over the age of 65 years may benefit from screening for the folate receptor autoantibody.

3. Targeting Transcobalamin-Receptor Pathway in cancer therapy: This ambitious multi pronged approach to targeting cancer cells uses a novel essential nutrient depletion strategy to slow or block cancer cells from proliferating. This strategy is likely to be less toxic and less likely to induce resistance to therapy. The monoclonal antibodies, the TC ligand and the receptor are also candidate carriers for delivering known chemotherapeutic drugs, toxins, imaging compounds and radionuclides to tumors. This project is being developed with the support of a Toronto based biotechnology company, KYTO Biopharma.

Research Interests:

My research interests include biochemical and molecular aspects of vitamin B12 absorption, transport and metabolism; Genetic abnormalities of vitamin B12 dependent pathways; Cellular and metabolic consequences of vitamin B12 and folate deficiencies; Vitamin B12, folate and homocysteine metabolism in the elderly population with cardio-vascular disease and cognitive disorders including Alzheimer's dementia; Neuropathology of vitamin B12 and folate deficiency; B12, folate status and DNA methylation in the brain. Another area of research actively being pursued in my laboratory is the association of folate receptor autoimmunity with neural tube defect pregnancy and cerebral folate deficiency.

Current research in my laboratory is focused on developing cancer therapeutics based on vitamin B12 and folate analogs that are anti-metabolites, drug-conjugates and pro-drugs; selective targeting of drugs via the B12 and folate transport proteins and their receptors; vitamin B12 and folate depletion strategies to inhibit cell replication in malignancies utilizing monoclonal antibodies to block cellular uptake of these vitamins and to deliver drugs. These approaches are designed to provide preferential targeting of rapidly dividing cancer cells because of the increased expression of receptors for uptake and increased need for these vitamins in cellular replication. In continuing our work on characterizing vitamin B12 binding proteins and the genes encoding these proteins, we are now studying the membrane receptor for the cellular uptake of vitamin B12. Having isolated the protein and gene encoding this receptor, we are now in a unique position to characterize this gene. It also provides us the tools for gene knockout studies that may finally yield valuable information on the role of vitamin B12 in embryogenesis, and the nervous system and perhaps identify the biochemical mechanism underlying the neuropathological manifestations of B12 deficiency which to date remains unexplained.

Miscellaneous Duties:

2012- Consultant, PAM Labs

2010 – Consultant, Par/Strativa Pharma

2009 – Present, Consultant, Covidien Inc.

2007- Present, Member, Executive Committee, SUNY Downstate Medical Center

2008-2010, Chairman, Research Resources Subcommittee

2007 – 2008 Consultant, IMMCO Diagnostics, Buffalo, NY.

1999 - Present, Consultant, KYTO Biopharma, Toronto, Canada.

1993-1997, Consultant and Collaborator, Receptagen Corporation, Seatle, WA; Department of Radiochemistry, University of Washington, Seatle, WA, the Biomedical Research Centre, University of British Columbia, Vancouver, Canada and The National Research Council, Ottawa, Canada.

1993 - 96, Member, Research and Development Committee.

V.A Medical Center, Brooklyn, NY 11209

1989 – 91, Chairman, Animal Research Subcommittee,

V.A. Medical Center, Brooklyn, NY 11209

1986 – 88, Member, Research and Development Committee,

V.A. Medical Center, Brooklyn, NY 11209

1986 – 88, Member, Animal Research Subcommittee,

V.A. Medical Center, Brooklyn. NY 11209

Prior Research Funding:

1991-1995, Study of Transcobalamin II, a vitamin B12 transport protein in the blood. V.A. Merit Review. Veterans Affairs. (Principal Investigator: E.V. Quadros)

Amount: \$726,845.

1993-1996, Receptor -Transcobalamin as a target for apoptosis and for delivery of macro molecules.

Receptagen Inc. Seatle, Washington. (Principal Investigator: E.V. Quadros)

Amount: \$263,000.

1994-1999, Kinetics of B12, Intrinsic Factor and other proteins. NIH. (S.P. Rothenberg, Principal Investigator; E.V.Quadros, CoInvestigator)

Amount: \$934,984.

1999 – 2000. Receptor-Transcobalamin II as a target for inducing apoptosis and for delivery of therapeutic compounds. BTwelve Inc. Toronto, CA. (Principal Investigator: E.V. Quadros,)

Amount: \$124,000

2000-01, Amount \$128,000

1999 - Axis Shield PLC, Oslo, Norway.

Monoclonal antibody based assay development.

1999 - 2000, Amount \$15,050.

2004-2005 – NIH (KO7), NIA Alzheimer's Disease Pilot Project.

Amount: **\$20,000**

April 2006 – March 2009. NIH R21 HD051880-01

Amount: \$ 166,000

July 2011, SUNY Technology accelerator award: \$50,000

Current Research Funding:

1. Autism Speaks June 2013 – May 2016 Basic and Clinical Research Grant Amount: \$449,00

2. Sep 2005 – June 2012. NIH RO1 DK064732-01A1 Amount: \$ 1,100,000 (Includes ARRA and bridge funding)

3. Nov. 2006 – multi year contract, KYTO Biopharma, Toronto, Canada. Amount: \$ 134,000 for 2009 - 13.

Patents

- 1. The use of monoclonal antibodies to human transcobalamin II to deprive cells of cobalamin. EU Patent # 0783526, Issued March, 2006.
- 2. Method for detecting the affinity of folate receptor autoantibodies US#7846672B2, 2010
- 3. Transcobalamin receptor polypeptides, nucleic acids, and modulators thereof, and related methods of use in modulating cell growth and treating cancer and cobalamin deficiency. PUPA/US60/790330.

Symposia Publications:

1. **Quadros, E.V.**, Jackson, B., Hoffbrand, A.V. and Linnell, J.C. (1979). Interconversion of cobalamins in human lymphocytes in vitro and the effect of nitrous oxide on synthesis of

- cobalamin coenzymes. Third European Symposium on Vitamin B12 and Intrinsic Factor. (Zagalak, B. and Friedrich, W. eds.), pp. 1045-1054.
- 2. **Quadros, E.V.** and Rothenberg, S.P. (1990). The structure and biosynthesis of human Transcobalamin II. Cobalamin '88 Proceedings of the First International Symposium on Biomedicine and Physiology of Vitamin B₁₂, p. 281-288, The Children's Medical Charity, London.
- 3. Rothenberg, S.P. and **Quadros, E.V**. (1990). Ligand binding and radioimmuno assay for cobalamin. Cobalamin '88 Proceedings of the First International Symposium on Biomedicine and Physiology of Vitamin B₁₂, p. 401-414, The Children's Medical Charity, London.

Book Chapters

- 1. Rothenberg, S.P. and **Quadros, E.V**. (1995). Transcobalamin II and the membrane receptor for the transcobalamin II-cobalamin complex. In Bailliere's Clinical Haematology (Wickramasinghe, S.N. ed) 8: 449-514.
- 2. Rothenberg, S.P., and **Quadros, E.V**. (1997). Quantitative methods for measurement of transcobalamin II (C.A.Wagner, Ed) Methods in Enzymology. 281: 261-268.
- 3. Quian, L., **Quadros, E.V.**, and Rothenberg, S.P. (1997). Molecular methods for analysis and expression of transcobalamin II. (C.A. Wagner, Ed) Methods in Enzymology. 281: 269-281.
- 4. Rothenberg, S.P., **Quadros, E.V**. and Regec, A. (1999). Transcobalamin II. In Vitamin B12 (Bannerjii, R. Ed). John Wiley and Sons Publ.
- 5. **Quadros, E.V**. Quantitative assay of vitamin B₁₂ In Modern Analytical Methods on Fat Water-Soluble Vitamins. (W.O. Song and G.R. Beecher Ed.) John Wiley & Sons Publ. (2001). p 313-328.
- 6. Ramaekers, V.T. and **Quadros, E.V.** (2010). Folate receptor autoimmunity in cerebral folate deficiency. (R.C. Dale and A Vincent Ed.) Clinics in Developmental Medicine, 184 185, Mac Kieth Press, London, UK.

Thesis Publications:

- 1. **Quadros, E.V**. (1972). Nutritional disorders of the nervous system: Studies on vitamin B₁₂, thiocyanate and folates. M.Sc. Thesis, University of Bombay, India.
- 2. **Quadros, E.V**. (1976). Studies on the distribution of cobalamins in man and animals and on cellular synthesis of cobalamin coenzymes in-vitro. Ph.D. Thesis, University of London (UK).

Conferences and meetings: (Abstracts deleted)

PEER REVIEWED PUBLICATIONS:

- 1. Dastur, D.K., **Quadros, E.V.**, Wadia, N.H., Desai, M.M. and Bharucha, E.P. (1972) Effect of vegetarianism and smoking on Vitamin B₁₂, thiocyanate and folate levels in the blood of normal subjects. Br. Med. J. 3:260-262.
- 2. Wadia, N.H., Desai, M.M., **Quadros, E.V**. and Dastur, D.K. (1972) Role of vegetarianism, smoking and hydroxocobalamin in optic neuritis. Br. Med. J. 3:264-267.
- 3. Dastur, D.K., Santha Devi, N., **Quadros, E.V.**, Gagrat, B.M., Wadia, N.H., Desai, M.M., Singhal, B.S. and Bharucha, E.P. (1975) Interrelationships between the B-vitamins in B₁₂ deficiency neuromyelopathy: A possible malabsorption-malnutrition syndrome. Am. J. Clin. Nutr. 28:255-270.
- 4. **Quadros, E.V.**, Matthews, D.M., Wise, L.J. and Linnell, J.C. (1975) Known and unknown cobalamins in the rat. Clin. Sci. Mol. Med. 48:4-5.
- 5. **Quadros, E.V.**, Matthews, D.M., Wise, L.J. and Linnell, J.C. (1976) Tissue distribution of endogenous cobalamins in the Rat, Cat and Guinea Pig. Biochim. Biophys. Acta 421:141-152.
- 6. Dastur, D.K., Santha Devi, N., **Quadros, E.V.**, Gagrat, B.M., Wadia, N.H., Desai, M.M. and Bharucha, E.P. (1976) B-vitamins in malnutrition alcoholism: interrelationships in blood and CSF. Br. J. Nutr. 36:143-159.
- 7. **Quadros, E.V.**, Matthews, D.M., Hoffbrand, A.V. and Linnell, J.C. (1976) Synthesis of cobalamin coenzymes by human lymphocytes in vitro and the effects of folates and metabolic inhibitors. Blood 48:609-619.
- 8. Linnell, J.C., **Quadros, E.V**., Matthews, D.M., Morris, H.P. and Poirier, L.A. (1977) Altered cobalamin distribution in rat hepatomas and in the livers of rats treated with diethylnitrosamine. Cancer Res. 37:2975-2978.
- 9. Hansen, O.P., Drivsholm, A., Hippe, E., **Quadros, E.V**. and Linnell, J.C. (1978) Interrelationships between vitamin B₁₂ and folic acid in myelomatosis: cobalamin coenzyme and tetrahydrofolic acid function. Scand. J. Haematol. 20:360-370.
- 10. **Quadros, E.V.**, Hamilton, A., Matthews, D.M. and Linnell, J.C. (1978) Isolation of ⁵⁷ Cocobalamin coenzymes at high specific activity from Streptomyces griseus. J. Chromatogr. 160:101-108.
- 11. Myasishcheva, N.V., **Quadros, E.V**., Matthews, D.M. and Linell, J.C. (1979) Interference by methylcobalamin analogues with synthesis of cobalamin coenzymes in Human Lymphocytes in vitro. Biochim. Biophys. Acta, 588:81-88.
- 12. Linnell, J.C., **Quadros, E.V**., Elliott, P.G. and Malleson, P. (1980) Defective adenosylcobalamin synthesis in a case of transcobalamin-II deficiency. J. Inher. Metab. Dis. 3:95-96.

- 13. Linnell, J.C., **Quadros, E.V**., England, J.M., Down, M.C. and Reynolds, E.G. (1981) Abnormal Cobalamin metabolism in a case of juvenile pernicious anemia with neurological symptoms. J. Inher. Metab. Dis. 4:149-150.
- 14. Jacobsen, D.W., Green, R., **Quadros, E.V**. and Montejano, Y.D. (1982) Rapid analysis of cobalamin coenzymes and related corrinoid analogs by high-performance liquid chromatography. Anal. Biochem. 120:394-403.
- 15. **Quadros, E.V.**, Rothenberg, S.P., Pan, Y. Ch and Stein, S. (1986) Purification and molecular characterization of human transcobalamin II. J. Biol. Chem. 261:15455-15460.
- 16. **Quadros, E.V.**, Rothenberg, S.P. and Polu, S. (1988) A specific radioimmunoassay for the measurement of 5' deoxyadenosyl cobalamin in serum. Brit. J. Haematol. 69:551-557.
- 17. **Quadros, E.V.**, Rothenberg, S.P. and Jaffe, E. (1989) Human endothelial cells synthesize and secrete functional transcobalamin II. Amer .J. Physiol. 256:C296-C303.
- 18. Platica, O., Janeczko, O., Regec, A, **Quadros, E.V**. and Rothenberg, S.P. (1989) Isolation of the complementary DNA for human transcobalamin II. Proc. Soc. Exptl. Biol. Med. 192:95-97.
- 19. Platica, O., Janeczko, R., **Quadros, E.V.**, Regec, A., Romain, R. and Rothenberg, S.P. (1991) The cDNA sequence and the deduced amino acid sequence of human transcobalamin II share homology with rat intrinsic factor and human transcobalamin I. J. Biol. Chem. 266:7860-7863.
- 20. **Quadros, E.V.**, Sai, P. and Rothenberg, S.P. (1993) Functional human transcobalamin II Isoprotein are secreted by insect cells using the baculovirus expression system. Blood. 81:1239-1245.
- 21. **Quadros, E.V.**, Sai, P. and Rothenberg, S.P. (1994) Characterization of the human placental membrane receptor for transcobalamin II- Cobalamin. Arch.Biochem.Biophys.308:192-199.
- 22. Regec, A. **Quadros, E.V.**, Platica, O. and Rothenberg, S.P. (1995) The cloning and characterization of the human transcobalamin II gene. Blood. 85: 2711-2719.
- 23. **Quadros, E.V**. and Jacobsen, D.W. (1995) The dynamics of cobalamin utilization in L-1210 mouse leukemia cells; a model of cellular cobalamin metabolism. Biochim. Biophys. Acta.1244: 395-403.
- 24. **Quadros, E.V.**, Rothenberg, S.P. and McLoughlin, P. (1996) Characterization of monoclonal antibodies to functional epitopes of human transcobalamin II. BBRC 222:149-154.
- 25. Pathare, P.M., Wilbur, S.D., Heusser, S., **Quadros, E.V.**, McLoughlin, P., and Morgan, A.C. (1996) Synthesis of cobalamin-biotin conjugates that vary in the position of coupling. Evaluation of cobalamin derivative binding to transcobalamin II. Bioconjugate Chemistry. 7:217-232.

- 26. Wilbur, S.D., Hamlin, D.K., Pathare, P.M., Heusser, S., Vessella, R.L., Buhler, K.R., Stray, J.E., Daniel, J., **Quadros, E.V.**, McLoughlin, P. and Morgan, A.C. (1996) Synthesis and nca-radioiodination of arylstannyl-cobalamin conjugates. Evaluation of aryliodo-cobalamin conjugate binding to transcobalamin II and biodistribution in mice. Bioconjugate Chemistry. 7: 461-474.
- 27. McLean, G.R., **Quadros, E.V.**, Rothenberg, S.P., Morgan, A.C., Schrader, J.W. and Ziltener, H.J. (1997) Antibodies to transcobalamin II block in-vitro proliferation of leukemic cells. Blood 89: 235-242.
- 28. Pathare, P.M., Wilbur, D.S., Hamlin, D.K., Heussea, S., **Quadros, E.V.**, McLoughlin, P., and Morgan, A.C. (1997) Synthesis of cobalamin dimers using isophthalate cross-linking of corrin ring carboxylates and evaluation of their binding to transcobalamin II. Bioconjugate Chem. 8: 161-172.
- 29. Wilbur, S.D., Pathare, P.M., Hamlin, D.K., Rothenberg, S.P. and **Quadros, E.V**. (1999) Radioiodination of cyanocobalamin conjugates containing hydrophobic linkers: Preparation of a radionated cyanocobalamin monomer and two dimers and assessment of their binding with transcobalamin II. Bioconjugate Chemistry. 10: 912-920.
- 30. **Quadros, E.V.**, Regec, A., Khan, K.M.F., Quadros, E. and Rothenberg, S.P. (1999) Transcobalamin synthesized in the intestinal villi facilitates transfer of cobalamin to the portal blood. Amer. J. Physiol. 277: G161-G166.
- 31. Sobti, P., Rothenberg, S.P. and **Quadros, E.V**. (2000) Radio-enzymatic assay for reductive catalysis of N⁵,N¹⁰-methyltetrahydrofolate by methylenetetrahydrofolate reductase. J. Biochem. Biophys. Methods 46: 11 20.
- 32. Ullend M, Eilertsen, I., **Quadros, E.V.,** Rothenberg, S.P., Fedosov, S.N., Sundrehagen, E. and Orning, L. (2002) Direct assay for cobalamin bound to transcobalamin (holotranscobalamin) in serum. Clin Chem 48:526 532.
- 33. Qian, L., **Quadros, E.V.**, Rejec, A., Zitouan, J. and Rothenberg, S.P. (2002) Congenital Transcobalamin II deficiency due to errors in RNA editing. Blood Cells Mol. Dis. 28: 134-142.
- 34. Regec, A., **Quadros, E.V.** and Rothenberg, S.P. (2002) Transcobalamin II expression is regulated by transcription factor(s) binding to a hexameric sequence (TGGTCC) in the promoter region of the gene. Arch. BioChem. Biophys. 407: 202-208.
- 35. Namour, F., Helfer, A-C., **Quadros, E.V**., Alberto, J-M., Bibi, Haim., Orning, L., Rosenblatt, D and Gueant, J-L. (2003) Transcobalamin deficiency due to activation of an intra exonic cryptic splice site. Brit. J. Haemat. 123: 915-920.
- 36. Gordon, M.M., Brada, N., Ramacha, A., Badell, I., delRio, E., Baiget, M., Santer, R., **Quadros, E.V.**, Rothenberg, S.P. and Alpers, D.H. (2004) A genetic polymorphism in the coding

- region of the gastric intrinsic factor gene is associated with congenital intrinsic factor deficiency. Hum. Mutat. 23: 85-91.
- 37 Fawwaz, Y., Rothenberg, S.P., Rao, S., Gordon, M.M., Alpers, D.H. and **Quadros, E.V**. (2004) Inherited deficiency of intrinsic factor due to a four base deletion in the gene. Blood. 103: 1515-1517.
- 38. Rothenberg, S.P., daCosta, M.P., Sequeira, J.M., Cracco, J., Roberts, J.L., Weedon, J. and **Quadros, E.V**. (2004) Autoantibodies against folate receptors in women with a pregnancy complicated by a neural tube defect. NE.JM. 350: 134-142.
- 39. **Quadros, E.V.**, Nakayama, Y., Sequeira, J.M. (2005) The binding properties of the human receptor for the cellular uptake of vitamin B12 Biochemical and Biophysical Research Communications 327 1006–1010
- 40. Ramaekers, V.Th., Rothenberg, S.P., Sequeira, J.M., Opladen, T., Blau, N., **Quadros, E.V.**, Selhub, J. (2005) Autoantibodies against folate receptors are associated with the infantile onset cerebral folate deficiency syndrome NEJM. 352, 1985–1991.
- 41. Fedosoy, S.N, Örning, L, Løvli, T,. **Quadros, E.V**., Thompson K, Berglund, L Petersen, T.E. (2005) Mapping the functional domains of human transcobalamin using monoclonal antibodies. FEBS J. 272, 3889 3898.
- 42. Orning, L., Rian, A., Campbell, A., Brady, J., Fedesov, S., Bramlage, B., Thompson, K and **Quadros, E.V**. (2006) Characterization of a monoclonal antibody with specificity for holo-transcobalamin. Nutr. Metab. (Lond). 3:3
- 43. Guéant, J-L., Chabi, N., Guéant-Rodriguez, R-M., Osvaldo, M., Debard, R., Payet, C., Lu, X., Villaume, C., Bronowicki, J-P., **Quadros**, **E.V**., Sanni, A., Amouzou, E., Bing, X., Chen, M., Anello, G., Bosco, P., Romano, A., Arrieta, H., Sanchez, B., Romano, A., Herbeth, B., Anwar, W., Namour, F. (2007) Environmental influence on the world-wide prevalence of a 776C>G variant in the transcobalamin gene (TCN2). J. Med. Genet. 44, 363 367.
- 44. Ramaekers, V.Th. and **Quadros, E.V.** Folate receptor autoimmunity in cerebral folate deficiency. In Inflammatory and Autoimmune Disorders of the Nervous System in Children Chapter 19, pp 302 316, Mac Keith Press, London, 2010
- 45. Ramaekers, V.Th., Sequeira J.M., Artuch R., Blau N., Temudo T., Ormazabal A., Pineda, M., Aracil A., Roelens F., Laccone F and **Quadros E.V.** (2007) Folate Receptor Autoantibodies and Cerebral Folate Deficiency In Rett Syndrome. Neuropediatrics 38: 179 183.
- 46. Garcia-Cazorla, A., **Quadros, EV**., Nascimento, A., Garcia-Silva, MT., Paz Briones PhD, Montoya, J., Ormazábal, A., Artuch, R., Sequeira, JM., Blau, N., Arenas, J., Pineda, M. and Ramaekers, VTh. (2007) Mitochondrial diseases associated with cerebral folate deficiency. Neurology 15:1360-1362.

- 47. Ramaekers, V.Th., Weis, J., Sequeira, J.M., **Quadros, E.V.** and Blau, N. (2007) Mitochondrial Complex I Encephalomyopathy and Cerebral Folate Deficiency. Neuropediatrics Neuropediatrics 38: 184 187.
- 48. Ramaekers, V.Th., Blau, N., Sequeira, J.M., Nassogne, MC and **Quadros, E.V.** (2007) Folate Receptor Autoimmunity and Cerebral Folate Deficiency in low-IQ Autism with Neurological Deficits Neuropediatrics, 38: 276 281.
- 49. Ramaekers, V.Th., Sequeira, J.M., Blau, N and **Quadros, E.V.** (2008) A milk-free diet down regulates folate receptor autoimmunity in cerebral folate deficiency syndrome. DMCN, 50: 346 352.
- 50. Bonkowsky JL, Ramaekers VT, **Quadros EV**, Lloyd M (2008). Progressive Encephalopathy in a Child With Cerebral Folate Deficiency Syndrome. J Child Neurol. 12:1460-3
- 51. Berrocal-Zaragoza MI, Fernandez-Ballart JD, Murphy MM, Cavallé-Busquets P, Sequeira JM, **Quadros EV** (2009). Association between blocking folate receptor autoantibodies and subfertility. Fertil Steril. 91:1518-1521.
- 52. Berrocal-Zaragoza MI, Murphy MM, Ceruelo S, **Quadros EV**, Sequeira JM, Fernandez-Ballart JD. (2009) High Milk Consumers Have an Increased Risk of Folate Receptor Blocking Autoantibody Production but This Does Not Affect Folate Status in Spanish Men and Women. J Nutr. 139:1037-1041.
- 53. **Quadros, EV.,** Nakayama, Y., and Sequeira, J.M. (2009) The protein and the gene encoding the receptor for the cellular uptake of transcobalamin bound cobalamin. Blood 113:186-192.
- 54. Ament AE, Li Z, Sturm AC, Perko JD, Lawson S, Masterson M, **Quadros EV**, Tanner SM (2009). Juvenile cobalamin deficiency in individuals of African ancestry is caused by a founder mutation in the intrinsic factor gene GIF.

 Br J Haematol. 144:622-624.
- 55. Fine EJ, Miller A, **Quadros EV**, Sequeira JM, Feinman RD (2009). Acetoacetate reduces growth and ATP concentration in cancer cell lines which over-express uncoupling protein 2. Cancer Cell Int. 9:14.
- 56. Molloy AM, **Quadros EV**, Sequeira JM, Troendle JF, Scott JM, Kirke PN, Mills JL (2009). Lack of association between folate-receptor autoantibodies and neural-tube defects. N Engl J Med. 361:152-60.
- 57. Hasselmann, O; Blau, N; Ramaekers, VT; **Quadros, EV**; Sequeira, JM; Weissert, M (2010). Cerebral folate deficiency and CNS inflammatory markers in Alpers disease. Mol.

Gen. Metab 99:58-61.

- 58. Pangilinan F, Mitchell A, VanderMeer J, Molloy AM, Troendle J, M Conley M, Kirke PN, Sutton M, Sequeira JM, **Quadros EV**, Scott JM, J L Mills JL, Brody LC (2010) Transcobalamin II receptor polymorphisms are associated with increased risk for neural tube defects. J Hum. Genet. 47: 677 685
- 59. **Quadros EV**, Nakayama Y, Sequeira JM (2010) Targeted Delivery of Saporin Toxin by Monoclonal Antibody to the Transcobalamin Receptor, TCblR/CD320. Mol Cancer Ther. PMID: 20858723
- 60. Jiang W, Nakayama Y, Sequeira JM, **Quadros EV** (2011) Characterizing monoclonal antibodies to antigenic domains of TCblR/CD320, the receptor for cellular uptake of transcobalamin-bound cobalamin. Drug Delivery. 18: 74-78.
- 61. Jiang W, Sequeira JM, Nakayama Y, Lai Sh-C, **Quadros EV** (2010) Characterization of the promoter region of TCblR/CD320 gene, the receptor for cellular uptake of transcobalamin-bound cobalamin. Gene; 466; 49–55
- 62. **Quadros EV** (2010) Advances in the understanding of cobalamin assimilation and metabolism.. Br J Haematol 148:195-204.
- 63. **Quadros EV**, Lai,S-C, Nakayama Y, . Sequeira, JM, Hannibal, L, Wang, S, Jacobsen, DW Fedosov, S, Wright, E, Gallagher, RC, Anastasio, N, Watkins, D, Rosenblatt, DS (2010) Positive Newborn Screen for Methylmalonic Aciduria identifies the First Mutation in TCblR/CD320, the Gene for Cellular Uptake of Transcobalamin-Bound Vitamin B₁₂. Hum Mut 31:1 6.
- 64. Lai SC, Nakayama Y, Sequeira JM, **Quadros EV**. (2011). Down-regulation of transcobalamin receptor TCblR/CD320 by siRNA inhibits cobalamin uptake and proliferation of cells in culture. Exp Cell Res. 317: 1603 1607.
- 65. **Quadros EV**, Nakayama Y, Sequeira JM. Targeted Delivery of Saporin Toxin by Monoclonal Antibody to the Transcobalamin Receptor, TCblR/CD320.Mol Cancer Ther. (2010). 11:3033-3040.
- 66. Arendt JFB, **Quadros EV**, Nexo E Soluble transcobalamin receptor, sCD320, is present in human serum and relates to serum cobalamin Establishment and validation of an ELISA. Clin Chem Lab Med. 2012 Dec 5;50(3):515-9. doi: 10.1515/CCLM.2011.810. PMID: 22149746

- 67. Frye RE, Sequeira JM, **Quadros EV**, James JS, Rossignol DA. (2013)Cerebral folate receptor autoantibodies in autism spectrum disorder. Molecular Psychiatry, 18: 369-381.
- 68. Ramaekers VT, **Quadros EV**, Sequeira JM. (2013). Role of folate receptor autoantibodies in infantile autism. Molecular Psychiatry, 18: 270-271.
- 69. Fernàndez-Roig S, Lai S-C, Murphy MM, Fernandez-Ballart J, **Quadros EV**. Vitamin B_{12} deficiency in the brain leads to DNA hypomethylation in the TCblR/CD320 knockout mouse. Nutr Metab (Lond). 2012 May 18;9:41. doi: 10.1186/1743-7075-9-41. PMID: 22607050
- 70. Ramaekers VT, Sequeira JM, **Quadros EV**. (2013). Clinical recognition and aspects of cerebral folate deficiencies. Clinical Chemistry and Laboratory medicine, 51: 545-554.
- 71. Sequeira JM, Ramaekers VT, **Quadros EV**. (2013). The diagnostic utility of folate receptor autoantibodies in blood. Clinical Chemistry and Laboratory medicine, 51: 497-511.
- 72. **Quadros, EV** and Sequeira, JM (2013). Cellular uptake of cobalamin: transcobalamin and the TCblR/CD320 receptor. Biochimie, 95: 1008-1018.
- 73. Lai, SCh, Sequeira, JM, Nakayama, Y, Wlodarczyk, B, Cabrera, RM, Finnell, RH, Bottiglieri, T, **Quadros, EV**. (2013). The Transcobalamin Receptor Knockout Mouse: A Model for Vitamin B₁₂ Deficiency In the Central Nervous System. FASEB J 27: 2468 2475.
- 74. **Quadros, EV**, Nakayama, Y and Sequeira, JM. Saporin conjugated monoclonal antibody to the transcobalamin receptor TCblR/CD320 is effective in targeting and destroying cancer cells. J Cancer Therapy (2013). 4: 1074 1081.
- 75. Jiang W, Nakayama Y, Sequeira JM, **Quadros EV**.(2013) Mapping the functional domains of TCblR/CD320, the receptor for cellular uptake of transcobalamin-bound cobalamin. FASEB J. 27: 2988 2994.
- 76. Ramaekers VT, Pinto E, Sequeira JM, Ansseau M, Philippe P, Boemer F, V Bourse V, **Quadros, EV.** Folinic acid for Schizophrenia due to Folate Receptor Autoimmunity. Hum Mol Gen. 2014 In Press
- 77. Berrocal-Zaragoza MI, Sequeira JM, Murphy MM, Fernandez-BallarT JD, Abdel Baki SG, Bergold PJ, **Quadros, EV**. Folate deficiency in rat pups during weaning produces learning and cognitive deficits. Brit J Nutr. 2014 In Prress