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## Professional Biography

**Michael Tweedle, PhD** is the Stefanie Spielman Professor of Cancer Imaging and Professor of Radiology and Biophysics at The Ohio State University College of Medicine and James Comprehensive Cancer Center, adjunct Professor in the Chemistry and Biochemistry Department in the School of Arts and Sciences, and Director of the Wright Center Molecular Imaging Agent Laboratory. His education was at Knox College (BS Chemistry), Rice University (PhD Physical Chemistry) and Stanford University (NIH Research Service Fellow). His experience in diagnostic imaging began as a research scientist at New England Nuclear, Inc. In 1981 he invented the first catalyst for production of cationic Tc(III) radiopharmaceutical kits, and he worked on early targeted molecular imaging (MI) agents using monoclonal antibodies at NEN/DuPont Pharmaceuticals. In 1986, working with E.R. Squibb and Sons, he invented and developed the novel first generation macrocyclic Gd-based MRI agent, ProHance<sup>TM</sup> (gadoteridol), that is used in over a million patients annually. As President and chief scientist at Bracco Research USA Inc, he and his team created two new Molecular Imaging agents: a first in class targeted theranostic for prostate and breast cancer (LuAMBA) and a first in class VEGF-R2 angiogenesis receptor targeted US agent now in Phase-II clinical trials (BR55). His current work includes targeted theranostics, molecular imaging agents and Optical Surgical Navigation agents. He sits on the editorial boards of Magnetic Resonance Imaging and Investigative Radiology, has served on Expert Councils at the US Pharmacopeia, the elected Boards of Directors and as Officer of four professional societies, academic Scientific Advisory Boards in molecular Imaging, on Scientific Advisory Boards of public and private pharmaceutical and chemical companies, and the Dean's Scientific Advisory Board of New York University's School of Science. He has authored over 150 publications, including 38 US patents, 25 pending patent applications, and 20 book chapters and reviews. In 2005 he was awarded The Harry Fischer Medal for Excellence in Contrast Media Research.



## Curriculum Vitae

### Dr. Michael F. Tweedle

|                              |   |                            |  |
|------------------------------|---|----------------------------|--|
| <b><u>Office Address</u></b> | The Ohio State University<br>720 Biomedical Research Tower<br>460 W. 12th St., Office 720<br>Columbus, OH 43210<br>614 247-4427 | <b><u>Home Address</u></b> | 40 N Parkview Ave<br>Bexley, OH 43209<br>609 937-0096 cell |
|------------------------------|---|----------------------------|--|

**Education**      1969 - 1971 Joliet Junior College  
1971 – 1973 BA      Knox College (Chemistry)  
1973 – 1977 PhD     Rice University (Physical Chemistry)

### Postgraduate Training and Fellowship Appointments

1977 – 1980      National Research Service Award Stanford University

### Awards and Honors

|             |   |
|-------------|---|
| 1977        | National Research Service Award (NIH), Stanford University, Palo Alto, CA                       |
| 1988        | Editor, Special Issue, Nuclear Medicine and Biology   |
| 1990 -      | Editorial Board, Magnetic Resonance Imaging   |
| 1998 -      | Editorial Board, Investigative Radiology  |
| 1998        | Editor, Special Issue, Investigative Radiology (1998, 1999, 2000, 2005, 2009)                   |
| 2003        | Elected Board of Directors, Society of Noninvasive Imaging in Drug Development                  |
| 2005        | Awarded the Harry Fischer Medal For Excellence in Contrast Media Research (CMR – admin by RSNA) |
| 2005, 2007  | Spedding Award Committee, Rare Earth Research Conference  |
| 2006        | Elected Vice Chairman of Society of Noninvasive Imaging in Drug Development                     |
| 2007        | Elected, Chairman of Society of Noninvasive Imaging in Drug Development                         |
| 2007        | Board of Directors of The Academy of Molecular Imaging  |
| 2007 - 2012 | Science Dean's Advisory Board, New York University  |
| 2007, 2008  | University of Arizona, ARIBI External Advisory Committee  |
| 2008        | Elected, Secretary and Executive Committee, Academy of Molecular Imaging                        |
| 2008-2011   | Board of Trustees, Radiotracer Clearing House   |
| 2008        | Most Notable person, RT-Image, Top 25 Most Influential in Radiology                             |
| 2008 – 2011 | Program Committee, World Molecular Imaging Congress   |
| 2009        | Co-Organizer, Imaging in 2020 meeting, Jackson Hole, WY, USA                                    |

### **Awards and Honors (continued)**

- 2009 – Awarded Stefanie Spielman Chair in Cancer Imaging, Department of Radiology and The James Comprehensive Cancer Center at the Ohio State University
- 2009 – 2011 Centre for Probe Development and Commercialization, McMaster University, Canada
- 2010 - 2013 American Board of Science in Nuclear Medicine
- 2010 Mentor of the Year, Wright Center for Innovation
- 2011 - 2013 Chairman, Awards Selection Committee, Contrast Media Research Congress
- 2011 – 2013 Officer and Board Member, World Molecular Imaging Society
- 2012 - Co-Chairman and founder, Optical Surgical Navigation Interest Group
- 2013 Distinguished Undergraduate Research Mentor, OSU
- 2011 - 2014 Board of Directors, Officer, World Molecular Imaging Society

### **Faculty Appointments**

- 2009- Professor, Department of Radiology (home), and Professor in Biophysics Program  
The Ohio State University College of Medicine  
Stefanie Spielman Chair in Cancer Imaging  
Radiology Department, The James Comprehensive Cancer Center  
Professor, Adjunct, Department of Chemistry and Biochemistry, College of Arts and Sciences, OSU
- 2013 – 2014 Entrepreneurial Scholar, Technology Transfer Department, OSU
- 2006 – 2009 Adjunct Associate Professor, Department of Radiology  
University of Pennsylvania Medical School

### **OSU Internal Faculty Committees**

- 2012 – Faculty Council OSUMC (Elected)
- 2013 - College Assembly OSUMC (Elected)
- 2010 - Pharmacy / formulary (OSU Wexner)
- 2010 - Promotion and Tenure (OSU Radiology)
- 2014 - Chairman, Promotion and Tenure Committee, Radiology
- 2010 - 2013 Contrast Agent use Guidelines (Radiology)
- 2010 – 2013 Safety & Risk Reduction Committee (Radiology)

### **Past Professional Appointments**

- 1980-1983 Sr. Research Chemist, Radiopharmaceuticals  
New England Nuclear and E.I. DuPont de Nemours
- 1983 -1984 Section head, Contrast Agents  
The Squibb Institute for Medical Research
- 1984 - 1987 Assistant Director, Contrast Agents Research Department  
The Squibb Institute for Medical Research
- 1987 – 1989 Director, Contrast Media Research Department

1989 – 1992 The Squibb Institute for Medical Research  
Director, Contrast Media Research Department  
Bristol-Myers Squibb Pharmaceutical Research Institute  
1992 – 1994 Director of Chemistry, Diagnostics Drug Discovery  
Division Bristol-Myers Squibb Pharmaceutical Research Institute  
1992 – 1994 Promotion Review Board, Drug Discovery  
Bristol-Myers Squibb Pharmaceutical Research Institute  
1994 - 2009 President and CEO, Bracco Research USA Inc

### **Memberships in Professional Societies**

American Chemical Society  
Society of Nuclear Medicine  
International Society for Magnetic Resonance in Medicine  
Radiological Society of North America  
World Molecular Imaging Society

### **Consulting**

2009 - 2014 Baker Sterchi Cowden and Rice, Attorneys at Law  
2007- 2009 Gerhson Lehman

### **Scientific Advisory Boards**

2009 - 2012 Marillion Pharmaceuticals  
2009 – 2012 Macrocyclics Inc  
2011 – 2012 Arroyo Biosciences Inc  
2010 – 2014 Empirion  
2015 - Elyton, LLC (pending)

### **Corporate Boards of Directors**

1994 - 2009 Bracco Research USA Inc (private)  
2011 – 2014 Novelos Therapeutics (public) (now Cellectar Biosciences)

### **Honorary Positions**

2012- 2014 OSU College of Medicine Technology Assessment Board  
(reviewer; twice selected for presentation)  
2009 –2014 The Stefanie Spielman Chair in Cancer Imaging  
2011 -2014 Gold Medal Awards Committee Chairman (CMR)

### **Conference Organizing Boards**

1985 – 2014 Contrast Media Research biannual meeting  
2002 - Tc and other metals, Padua Italy  
1988 Antibody and MRI agent Chemistry  
2007, 2008 Rare Earth Research Society

### **Conference Organizing Boards (continued)**

|             |  |
|-------------|--|
| 2009        | co-Chairman Imaging in 2020                          |
| 2008 – 2010 | World Molecular Imaging Congress (Program Committee) |
| 2008 – 2010 | Therapy Emphasis Co-Chair,                           |
| 2010 - 2011 | Plenary Session Co-Moderator, WMIC                   |
| 2011        | Young Investigator Competition, WMIC                 |

### **Academic Boards (elected:E; appointed:A)**

|               |   |
|---------------|---|
| 2000 – 2005   | USP Expert Committees on Radiopharmaceuticals and Medical Imaging Agents (A)              |
| 2003 – 2006   | Board of Directors of Society of Noninvasive Imaging in Drug Development (E)              |
| 2005 - 2006   | Newsletter Editor, SNIDD (A)  |
| 2006 – 2007   | Vice-Chairman (Vice President) of SNIDD (E)   |
| 2007 – 2009   | Chairman (President) of SNIDD (E)   |
| 2007 - 2011   | Academy of Molecular Imaging (E)  |
| 2008 - 2011   | Executive Committee & Secretary (E)   |
| 2010          | Search Committee: Editor in Chief of Molecular Imaging in Biology Journal (A)             |
| 2006 – 2008   | External Scientific Grant Advisor (Arizona Cancer Center) (A)                             |
| 2007 - 2011   | Science Dean's Board of Advisor's, New York University (A)                                |
| 2008 – 2010   | NYU Science Dean Board Chair, Technology Transfer and Intellectual property Committee (A) |
| 2008 – 2010   | Radiotracer Clearing House, Board of Trustees (A)   |
| 2009 – 2011   | Centre for Probe Development and Commercialization, McMaster University, Canada 2010 (A)  |
| 2010 – 2013   | Board of Directors of the American Board of Science in Nuclear Medicine (A)               |
| 2010 - 2011   | Society for Molecular Imaging Council Member (A)  |
| 2011 - 2014   | World Molecular Imaging Society, Board of Trustees (A) World                              |
| 2011- 2014    | World Molecular Imaging Society Officer (Secretary) (E)                                   |
| <b>2015 –</b> | <b>Board of Directors, International Society of Image Guided Surgery (ISIGS)</b>          |

### **Editorial Positions**

|             |  |
|-------------|--|
| 1990 –      | Editorial Board of Magnetic Resonance Imaging                  |
| 1998 –      | Editorial Board of Investigative Radiology                     |
| 1988        | Editor, Special Issue, Nuclear Medicine and Biology            |
| 1998 - 2000 | Editor Special Issue, Investigative Radiology, Contrast Agents |
| 2005,       | Editor Special Issue, Investigative Radiology, Contrast Agents |
| 2010        | Editor Special Issue Investigative Radiology Contrast Agents   |
| 1990 – 2013 | J. of Magnetic Resonance in Medicine (reviewer)                |
| 1990 –      | Inorganic Chemistry (reviewer)                                 |
| 2001 –      | J. Am. Chem. Soc. (reviewer)                                   |
| 1991 –      | Radiology (consultant and Reviewer)                            |

- 1984 – Nuclear Medicine and Biology (reviewer)  
2013 - Molecular Imaging and Biology (reviewer)  
2011 – reviewer: PLOS One, Bioconjugate Chemistry, Inorganic Chemistry, J. Am. Chem. Soc., Org. Med. Chem. Lett., Acta Radiologica, Radiology, Mol. Imag. Biol.  
2009 – Cancer and imaging research (abstract reviewer)  
Abstract Reviewer: American Association Cancer Research, International Society of Magnetic Resonance in Medicine, Society for Noninvasive Imaging in Drug Development, Academy of Molecular Imaging, World Molecular Imaging Society  
2006 – 2007 Editor, Newsletter of the Society of Noninvasive Imaging and Drug Development (2006-7)

### Expert Panels and Study Sections

- 1990, 2009 NIH ad hoc Study Section: Program Project Grants and site visit  
1991 NIH ad hoc grant Study Section: Tissue Specific Contrast Agents  
1993 Diagnostic Imaging Research Branch Strategic Planning Advisor  
2002 Tc in Chemistry in Nuclear Medicine, Padua, Italy 2002  
2003 ISMRM Conf. on MRI in Cancer  
2004 NCI panel on (DCE) MRI in Cancer  
2005 NCI Conference on Diagnostics in Pharmaceutical Development, Bethesda, MD  
2005 Biochemical Imaging Agent Issues, International Society for Strategic Studies in Radiology  
2005 Radiology: Creating strategies for the next decade  
2008 Biomarkers Roundtable, by RSNA  
2009 Imaging Biomarkers Workshop, AMI/SNIDD  
2011 NIH NIBIB Ad Hoc Study Section S10 Grants  
2012 NIH, NIBIB Ad Hoc Study Section P41 Grant and site visit  
2011 – 2013 Denman Undergraduate Research Forum Faculty Judge  
2012 OSU Molecular Life Sciences Interdisciplinary Graduate Programs Symposium Poster Judge  
**2014 “Why are there no Molecular MRI agents?” International Society of Magnetic Resonance Imaging, Milan, Italy**  
**2015 NIH, Ad Hoc Study Section P41 Grant and Site visit**  
**2015 - DOD, CM Breast Cancer Research Program study section**

### Academic Funding

**Drug Development Institute, James Cancer Center. Role: PI, Co-PI Josh Goldberger. “Molecular Magnetic Resonance Imaging Probes that Self-Assemble in Malignant Tumors” The \$307,000 (\$185,000 year 1, with evaluations quarterly) bridges and augments an R21 being re-submitted July, 2015, that targets early breast cancer and many others. It’s aim is to generate the data required for a successful bid to venture capitalists from a start up company. It includes manufacturing drug product. Year 2 (funding ND)**

Drug Development Institute, James Cancer Center. “Localized delivery of Peptide Receptor Radionuclide Therapy” (i.a.PRRT) for focal therapy of early prostate cancer without generating side effects, designed to work with Active Surveillance to avoid whole gland removal. \$186,000 (1 year, evaluations quarterly). It’s aim is to generate the data required for a successful bid to venture capitalists from a start up company.

Pelotonia Idea Grant, OSU. Role: co-PI with Charles Hitchcock, “Identification and development of new immunoagents for cancer diagnosis and therapy.” 10/15 – 10/17, \$100,000

Pelotonia Idea Grant, OSU. Co-PI with Ron Xu and A. Yilmaz “Wearable Navigation System for Image-guided Cancer Resection Surgery” \$100,000, 10/14 – 10/16

2-U44-CA176664-02. NIH SBIR Phase II Contract. Role: Co-I. PI: M. V. Knopp; “Improved Metallofullerene MRI Contrast Agent for Cancer Diagnosis.” –. \$382,206. 2013-12/15

Role: PI. Enlyton Pharmaceuticals An assessment of the tumor-targeting capability of Enlyton’s antibody technology for use in PET imaging of cancer” \$189,000. 10/12 – 6/15

TECH13-060 N043CO-2012-00059. Ohio Third Frontier Innovation Platform Grant. IPP-12-254. Role: Co-PI with M. V. Knopp and K. Kumar, “Next Generation Multi-Modal Molecular Imaging Technology Platform”. \$6,026,690. 2013 – 2018.

SBIR 2447-NIH-2S/OSU (sub) to 2 U44 CA176664-02 NIH SBIR Phase II Role: Co-I. “Improved metallofullerene MRI contrast agent for cancer diagnosis.” PI; M. Knopp.

Pending Submissions (\$16,021,679)

RO1 to NCI/NIBIB RFA 13-185, Image-guided drug delivery in cancer (\$1,980,000, 4 years), to be submitted June 16, 2015, Role: PI, Co-PI: Thomas Rosol, Michael Knopp. “Image-guided, transcatheter Peptide Receptor Radiotheranostic for Prostate Cancer”

RO1 Grant GRT00039093. NIH. Co-I with PI: Miryam Lustberg, “Affective consequences of chemotherapy in breast cancer survivors”

R21 Grant 11679996. NIH. Role: Co-PI with J. Goldberger. “Molecular Magnetic Resonance Imaging Probes that Self-Assemble in Malignant Tumors”

R21 Grant GRT00038741. NIH. Co-PI with Chadwick Wright. “Innovative dual-function agent for image-guided cancer therapy.”

SSTR Phase I. GRT00037702. NIH. PI: M. F. Tweedle. Sponsor Enlyton, LLC. “Creation and preclinical development of a real time, wide field Optical Surgical Navigation imaging agent specific for TAG-72+ adenocarcinomas.”

**DOD Grant GRT00035750. DOD. Role: Co-PI with Ron Xu and A. Yilmaz. “Intraoperative integrated navigation for targeted breast-conserving surgery”**

**RO1 grant GRT00037727. NIH Role: Co-PI. PI: Ron Xu. “Multifunctional drug-loaded activatable microbubbles for ultrasound mediated combination therapy in ovarian cancer.”**

**RO1 grant. GRT00038811. NIH. Role: Co-I and PI on OSU subcontract. PI: James Basilion. “Theranostic gold nanoparticles for imaged-guided radical prostatectomy and PDT ablation.”**

**RO1 grant GRT00038338. NIH. Role: Co-I. PI: Dario Palmieri. “A novel class of anti-Nucleolin immunoagents for cancer therapy.”**

**RO1 grant GRT00038255. NIH. Role: Co-I. PI: Tonya Orchard. “Effect of omega 3 fatty acid supplementation and added sugars in the diet on chemotherapy-induced cognitive deficits.”**

**Enlyton Pharmaceuticals An assessment of the tumor-targeting capability of Enlyton's antibody technology for use in PET imaging of cancer” Role: PI. ~ \$300,000 – 400,000. For roll up to IND submittal**

### **Completed Funding**

9872-12: National Collegiate Inventors Association. PI: M. Tweedle with Undergraduates, K. Mitra and B. Geiger. “Bringing the Cancer Detection Kit Home” Phase II, \$25,000. 2014 - 2015

Bremmer CCTS, PI Chawick Wright. “<sup>123</sup>Iodine-labeled HN1-800 dye agents for dual-modality intraoperative surgical guidance during resection of head & neck squamous cell carcinoma” \$70,000. 2013 - 2014

Wright Center Molecular Imaging Agents Laboratory (OSU, OSUMC, Radiology): “Create an IP protected Drug” OSU, OSUMC, WCI and Radiology, \$5,200,000 2009 – 2014

Enlyton Pharmaceuticals. Contract PI, SBIR Contract No. HHSN261201200059C: “A Tunable Tumor Targeting Agent for SPECT Imaging” \$66,366. 5/20/12 – 6/30/15.

N043CO-2012-00059: “A tunable agent for SPECT Imaging.” PI: M. Tweedle \$50,607. Enlyton Inc. SBIR Phase I Contract 2013 – 2014

Pelotonia Idea Grant, PI: M. Tweedle, J. Goldberger, “Pan-cancer MRI agents that self-assemble in malignant tumors” \$100,000. 2012 – 2014

1R21CA159077 – 01. NCI. Co-PI with R. Xu, T Huang, R Shen, Radiology, Bioengineering, Exp. Genetics. “Multifunctional microbubbles for image-guided epigenetic therapy in cancer”. 2012-2014. \$421,000

RISE Award. James Comprehensive Cancer Center. “Dosimetry modulated radiotherapy with  $^{177}\text{Lu}$ AMBA, a new breast cancer theranostic.” Project V, Co-PI with J. White for a NCI Breast Cancer Spore Application. \$7,000.

CCTS Pilot Award. Co-PI with J. Kuret. Cancer Center Technical Services, “Tauopathic Imaging Agents.” \$25,000 2012

### **Lectures by Invitation (2008 – 2014)**

Status of AMI’s Molecular Imaging projects; RSNA sponsored round table on Molecular Imaging, Chicago IL, April, 2008

Degenerative Diseases, a Challenge for Imaging Solutions  
Italian Radiological Society Meeting; Rome, May 2008

Lu-AMBA, a  $^{177}\text{Lu}$ -containing peptide conjugate for simultaneous imaging and radiotherapy  
Rare Earth Research Conference; Tuscalusa, Alabama, June 2008

Lu-AMBA, a  $^{177}\text{Lu}$ -containing peptide conjugate for simultaneous imaging and radiotherapy  
International Macrocyclic Chemistry; Las Vegas, NV, July 2008

“Use of Multivalency in Drug Discovery in Molecular Imaging”  
Metabolism in Cancer Lecture Series; University of Pennsylvania, October 2008

"Peptide Targeted Molecular Imaging Agents"  
Molecular Imaging Seminar series; Stanford University, January 2009

“Matching the drugs to the patients”  
A Christmas to Cure Cancer sponsored by the Stefanie Spielman Foundation  
Columbus, OH, December 2009

“Cancer Discovery”  
Spielman Cruise for Cancer, Columbus, OH, February, 2010.

“M-AMBA, a novel thernostic for breast and prostate cancer”  
Imaging Research Seminar, Radiology Dept.; The Ohio State University, March, 2010

“The future of cancer detection and treatment through imaging technology”  
Stefanie’s Champions Breast Cancer Research Symposium  
Greater Columbus Convention Center, April 2010

"Comparative effectiveness of Contrast and Molecular Imaging agents"

Moffet Cancer Center; Tampa, Florida, May, 2010

“The future of cancer detection and treatment through imaging technology”  
Stefanie’s Champions Breast Cancer Research Symposium  
The Ohio Union, Columbus, OH April 2011

“Cancer Imaging: Past, Present and Future”  
Columbus Rotary Luncheon Meeting; Hyatt Regency Columbus, April 2011

“Molecular Imaging Agents: the Can, the Can’t and the Won’t”  
Cooley Visiting Professor Lecture, Radiology Dept.; University of Texas Medical Branch,  
November 2011

**“Can MRI Agents Compete in the Biochemical Arena?”**  
**National American Chemistry Society Meeting; Dallas Tx, March 2014**

**“Optical Surgical Navigation Probes”**  
**Cardinal Health Inc.; April, 2014, Dublin, Ohio**

**“Molecular Imaging MRI Probes that Self-Assemble in Malignant Tumors”**  
**Bracco Group; Milan, Italy, May 2014**

**“Clinical Applications, Workflow and Regulatory Issues: Intraoperative Guided  
Surgery with  
Optical Probes” 9-14, at WMIC, Seoul, Korea.**

#### **Internal OSU Teaching (2014)**

"Molecular Imaging Agents 101". Department of Radiology Residents' Seminar OSU presented annually.

“The Basic Principles and History of X ray and MRI Contrast Agents.”  
Department of Radiology Residents' Seminar, OSU Presented annually.  
“Recent Research”

“Molecular Imaging Research” Department of Radiology Residents' Seminar, OSU presented bi-annually.

Individual Studies. The Ohio State University. 2014 GRA Christian Beutner (Chemistry and Biochemistry) Primary Instructor

“Preclinical Analysis of infrared fluorescence labeled Probe to target bombesin receptors”  
(poster at James CCC annual Event)

Research in Biophysics. The Ohio State University. Primary Instructor:

**PhD Candidacy Examination. Xiaoli Liu, Biophysics. 11-24-14**

**PhD Oral Examination. Sara Lim. Biophysics.11-6-14.**

**Biophysics Rotation, Keyton Clayson. Biophysics. Nov. - Dec, 2014.**

**Xu Zhang. NSF I-Corps support. Bioengineering, 11-14**

**PhD. Pre-Oral. Maria Isabella Menendez. 12-17-14.**

**PhD. Oral Examination. Maria Isabella Menendez. June 16, 2015**

## **Bibliography**

**Patents** (only US Patent and published Applications listed)

1. M. F. Tweedle. Accelerators for Forming Cationic Tc Complexes Useful in Kits for Radiodiagnostic Heart and Hepatobiliary Tissue Imaging.  
US Patent Number 4,455,291, 1984
2. M.F. Tweedle, G.T. Gaughan, J.H. Hagan. Substituted Triscarboxymethyl-tetraazacyclododecane and Analogs  
US Patent Number 4,885,363, 1989
3. M.F. Tweedle, L.D. Bradshaw, L. J. Wilson. Paramagnetic Metalloporphyrins as Contrast Agents for Magnetic Resonance Imaging  
US Patent Number 5,262,532, 1993
4. J.F. Desreux, M.F. Tweedle, P.C. Ratsep, T.R. Wagler, E.R. Marinelli. Hepatobiliary Tetraazamacrocyclic Magnetic Resonance Contrast Agents  
US Patent Number 5,358,704, 1994
5. M.F. Tweedle, H.W. Strauss, A.D. Nunn. Methods for the In Vivo Measurement of the Concentration of non-Imaging NMR-Detectable Xenobiotic Compounds  
US Patent Number 5,468,467, 1995
6. M.F. Tweedle, G.T. Gaughan, J.J. Hagan. Method for Imaging Mammalian Tissue using 1-Substituted-1,4,7-Tricarboxy-Methyl-1,4,7,10-Tetraazacyclo-Dodecane and Analogs  
US Patent Number 5,474,756, 1995
7. R.S. Ranganathan, E. Marrinelli, R. Pilli, M. F. Tweedle. Aromatic Amide Compounds and Metal Chelates Thereof for Diagnostic Imaging  
US Patent Number 5,573,752, 1996
8. M.F. Tweedle, H.W. Strauss, A.D. Nunn  
Methods for the In Vivo Measurement of the Concentration of NMR-Detectable Xenobiotic Compounds  
US Patent Number 5,603,917, 1997

9. M.F. Tweedle, G.T. Gaughan, J.J. Hagan  
Method for Imaging Mammalian Tissue using 1-Substituted-1,4,7-Tricarboxy-Methyl-1,4,7,10-Tetraazacyclo-Dodecane and Analogs  
US Patent Number 5,674,470, 1997
  
10. M.F. Tweedle, G.T. Gaughan, J.J. Hagan  
Method for Imaging Mammalian Tissue using 1-Substituted-1,4,7-Tricarboxy-Methyl-1,4,7,10-Tetraazacyclo-Dodecane and Analogs  
US Patent Number 5,846,519, 1998
  
11. R. Ranganathan, M.F. Tweedle, P. Wedeking. Methods and Compositions for Using Non-ionic Contrast Agents to Reduce the Risk of Clot Formation in Diagnostic Procedures  
US Patent Number 5,869,024, 1999
  
12. R. Ranganathan, T. Arunachalam, M.F. Tweedle. N, N-Dimethyldiatrizoic Acid and its conjugates as Hepatobiliary agents for X ray CT Imaging  
US Patent Number 6,051,210, 2000
  
13. J. Desreux, P. Jaques, P. Humblet, P. Herman, P. Comblin, P. Tholet, M. F. Tweedle. Self Assembling Heteropolymetallic Chelates As Imaging Agents and Radiopharmaceuticals  
US Patent Number 6,056,939, 2000
  
14. P. Wedeking, T R. Wager, T. Arunachalam, R. Ramalingam, K. Linder, R. Ranganathan, A. Nunn, K. Raju, M.F. Tweedle. Metal Complexes Derivatized with Folate for use Diagnostic and Therapeutic Applications  
US Patent Number 6,093,382, 2000
  
15. M. Tweedle, G. Gaughan, J. Hagan. Method for Imaging and Radiopharmaceutical Therapy using 1-substituted-4,7,10-tricarboxymethyl-1,4,7,10-tetraazacyclododecane and analogs  
US Patent Number 6,143,274, 2000
  
16. P. Wedeking, T R. Wager, T. Arunachalam, R. Ramalingam, K. Linder, R. Ranganathan, A. Nunn, K. Raju, M.F. Tweedle. Metal Complexes Derivatized with Folate for use Diagnostic and Therapeutic Applications  
US Patent Number 6,221,334, 2001
  
17. R. Ranganathan, T. Arunachalam, M.F. Tweedle. N, N-Dimethyldiatrizoic Acid and its conjugates as Hepatobiliary agents for X ray CT Imaging  
US Patent Number 6,264,916, 2001
  
18. R. Ranganathan, X. Zhang, R. Shukla, M.F. Tweedle. Enhanced Relaxivity Monomeric and Multimeric Compounds  
US Patent Number 6,693,190, 2004

19. R. Pillai, S. I. Kang, E. Marinelli, R. Ranganathan, M.F. Tweedle. "Aminocarboxylate ligands having substituted aromatic amide moieties"  
US Patent Number 6,875,864, 2005
20. M. A. Von Wronski, E.R. Marinelli, A. D. Nunn, R. Pillai, K. Ramalingam, M.F. Tweedle, K. Linder, P. Nanjappan, N. Raju. Compounds for targeting endothelial cells, compositions for the same and methods for their use  
US Patent Number 7,109,167, 2006
21. R. Ranganathan, H. Fan, M.F. Tweedle. Conjugates of antioxidants with metal chelating ligands for use in diagnostic and therapeutic applications  
US Patent Number 7,160,535, 2007
22. P. Wedeking, T R. Wager, T. Arunachalam, R. Ramalingam, K. Linder, R. Ranganathan, A. Nunn, K. Raju, M.F. Tweedle. Metal Complexes Derivatized with Folate for use Diagnostic and Therapeutic Applications  
US Patent Number 7,186,397, 2007
23. E. Cappelletti, L. Lattuada, K.E. Linder, E. Marinelli, P. Nanjappan, N. Raju, R. Swenson, M.F. Tweedle. Gastrin Releasing Peptide Compounds  
US Patent Number 7, 226,577, 2007
24. R. Pillai, S. I. Kang, E. Marinelli, R. Ranganathan, M.F. Tweedle. Aromatic Amide Compounds and Metal Chelates Thereof  
US Patent Number 7,351,389, 2008
25. C. A. Chang; K. Kumar, M.F. Tweedle. Dual functioning excipient for metal chelate contrast agents.  
US Patent Number 7,385,041, 2008
26. P. Wedeking, T R. Wager, T. Arunachalan, R. Ramalingam, K. Linder, R. Ranganathan, A. Nunn, K. Raju, M.F. Tweedle. Metal Complexes Derivatized with Folate for use Diagnostic and Therapeutic Applications  
US Patent Number 7,399,460, 2008
27. R. Ranganathan, H. Fan, M.F. Tweedle. Conjugates of Antioxidants with Metal Chelating Ligands for use in Diagnostic and Therapeutic Applications  
US Patent Number 7,407,644, 2008
28. R. Ranganathan, H. Fan, M.F. Tweedle, R.E. Swenson. Conjugates of Antioxidants with Metal Chelating Ligands for use in Diagnostic and Therapeutic Applications.  
US Patent Number 7,582,280, 2009
29. E. Cappelletti, L. Lattuada, K.E. Linder, E. Marinelli, P. Nanjappan, N. Raju, K. Ramalingam, R. Swenson, M.F. Tweedle. Gastrin Releasing Peptide Compounds  
US Patent Number 7,611,692, 2009

30. M.A. Von Wronski, E.R. Marinelli, A.D. Nunn, R.K. Pillai, K. Ramalingam, M.F. Tweedle, K.E. Linder, P. Nanjappan, N. Raju, F. Yan, M. Schneider. Compounds for targeting endothelial cells, compositions containing the same and methods for their use  
US Patent Number 7,820,621, 2010

31. E. Cappelletti, L. Lattuada, K.E. Linder, E. Marinelli, P. Nanjappan, A.D. Nunn, N. Raju, K. Ramalingam, R.E. Swenson, M.F. Tweedle. Gastrin releasing peptide compounds  
US Patent Number 7,850,947, 2010

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**Recent Abstracts (2009 – 2014) 3 new in 2013/14 (Only External Abstracts included)**

- 1.. K.E. Linder, E. Metcalfe, P. Nanjappan, T. Arunachalam, T. Skedzielewski, K. Ramos, A.D. Nunn, M.F. Tweedle, R.E. Swenson "Black Hole Quencher 3 (BHQ-3) quenches fluorescence from IRDye800 but is unstable in vivo - a tale of two compounds." World Molecular Imaging Congress, Montreal, Canada, September 2009
- 2.. K.E. Linder, E. Metcalfe, P. Nanjappan, T. Arunachalam, T. Skedzielewski, K. Ramos, A.D. Nunn, M.F. Tweedle, R.E. Swenson. "Light imaging agents containing IRDye800 are contact quenched by Black Hole Quencher 3" Contrast Media Research 2009. Copenhagen, Denmark, October 2009

- 3.. R. Xu, J.S. Xu, J. Huang, M.F. Tweedle, C.Schmidt, S.P. Povoski, E.W. Martin  
“Targeted delivery of cancer-specific multimodal contrast agents for intraoperative detection of tumor boundaries and therapeutic margins” Design and Quality for Biomedical Technologies III. San Francisco, California, January 2010
4. R. Layman, N.C. Hall, J. Zhang, D. Barker, M. Natwa, M.F. Tweedle, M.V. Knopp.  
“Overview of current technology in clinical hybrid SPECT/CT” Society of Nuclear Medicine 57<sup>th</sup> Annual Meeting 2010. Scientific Poster Session: Educational Exhibit. Salt Lake City, Utah, June 2010
5. J. D. Clifton, N. Beardsley, K.E. Linder, P. Nanjappan, T. Mawn, E. Metcalfe, R. Eisenberg, M.F. Tweedle, J. Delikatny. “Non invasive in-vivo optical imaging of matrix metalloproteinase activity in a KRN serum transfer model of Arthitic mice with novel near infra-red probes.” University of Pennsylvania 9<sup>th</sup> Annual Biomedical Postdoctoral Research Symposium. Philadelphia, Pennsylvania, October 2010
6. Jeff Xu, Jiwei Huang, Ruogu Qin, Michael Tweedle, Ronald Xu. 2010. “Gold Nanoparticle Assisted Light Activation of Microbubbles for Photothermal Therapy Photonics West” 2010: 7551-32.
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8. K. Kumar, M. Matson, L.J. Wilson, M.F. Tweedle. “Loading and Human Plasma Stability of Gadonanotubes” World Molecular Imaging Congress, San Diego, California, September 2011
9. S.H. Wang, N. Raju, H. Ding, M.F. Tweedle. “Evaluation of biotinylated Bombesin analogs as tumor targeting probes” World Molecular Imaging Congress, San Diego, California, September 2011
- 10 K. Kumar, M. Matson, L.J. Wilson, M.F. Tweedle. “On the Stability of Gadonanotubes” Contrast Media Research Conference” Galveston, Texas, October 2011
11. Q. Ma, M.F. Tweedle. “X-ray Absorption Spectroscopy Study of Gd<sup>3+</sup> loaded ultra short carbon nanotubes” 15<sup>th</sup> International Conference on X-ray Absorption Fine Structure (XAFS-XV), Beijing, China July 2012
12. Shrivastava A, Wang S-H, Natarajan, R, Gierach I, Tweedle M, “Heterobivalent chelating ligand for targeting GRP and NPY-1 receptors on tumor cells” World Molecular Imaging Society, Dublin Ireland, September, 2012
13. Mohd Yusof A, Kothandaraman S, Ding H, Tweedle M, Phay J. “DEVELOPMENT OF A CALCIUM-SENSING RECEPTOR MOLECULAR IMAGING AGENT”.Annual AAES Meeting

Chicago, April 14 –16, 2013

14. A. Shrivastava, H. Ding, S. Kothandaraman, S. Wang, R. Natarajan, M. Williams, N. Raju, L. Gong, S. Zheng, K. Milum, M. F. Tweedle. “Near infrared fluorescent (NIRF) probes targeting bombesin receptors on breast and prostate cancers” Contrast Media Research 2013, November 2013. Beijing, China

15. C. Wright, R. Layman, M. F. Tweedle. “Optical Imaging of positron-emitting radionuclides using Cerenkov luminescence and its potential applications in radioguided surgery. Nafki , 2013

16. A. Ghosh, C. Beutner, M. Williams, H. Ding, A. Bratasz, K. Kumar, M. Tweedle, J. Goldberger. “Developing Pan Cancer Targeted MRI Contrast Agents that Self-Assemble in Malignant Tumors” ISMRM, Milan Italy, May, 2014

17. K. A. Collier, J. Zhang, K. Binzel, N. C. Hall, K. Kumar, M. F. Tweedle, P. Monroe, M. V. Knopp. “Neuroimaging PET tracers: An atlas based approach for comparison in NHP PET/CT” SNM 2014, St. Louis, Mo.

18. Haiming Ding, Shankaran Kothandaraman , Li Gong, Michelle M. Williams, Wessel P. Dirksen, Thomas J. Rosol, Michael F. Tweedle. “NIRF Peptides Targeted to HuGRPr-transfected Canine ACE-1 Prostate Cancer Tumors.” 2014 World Molecular Imaging Congress, Soel, Korea.

19. **Haiming Ding, william T. Drost, Li Gong, Shankaran Kothandaraman, Maria Menendez, Huyen nguyen, mitch phelps, Joshua Dowell, Robert R. Bahnson, Michael V. Knopp, Thomas J. Rosol, Michael F. Tweedle, “Orthotopic canine prostate cancer model for imaging agents to human cancer receptors” 2015 World Molecular Imaging Congress, Honolulu, Hawaii.**

20. **Chadwick L. Wright, Ricky R. Layman, Shankaran Kothandaraman, Nathan C. Hall, Michael V. Knopp, Michael F. Tweedle, “Buckeye-Boost: Cerenkov luminescence amplification by the wavelength-shifting Ohio Buckeye extract, esculin.” ACNM, 2015**

21. **Ashley Braddom, Timothy Richmond, Tyler Sheetz, Erika Reese, Anna Tessari, Kathleen Tober, Christin E. Burd, Claudia De Lorenzo, Edward W. Martin, Jr., Vincenzo Coppola, Charles Hitchcock, Michael F. Tweedle, Tatiana Oberyszyn, Carlo M. Croce and Dario Palmieri. “Anti-NCL scFv as a new tool for skin cancer diagnosis and therapy.” James CCC Translational Therapeutics retreat. Oct 31, 2015, Columbus, Ohio**

22. **Tweedle, M.F. Dowell, J.; Bahnson, R.R.; Wright, C.L.; Ding, H.; Gong, L.; Kothandaraman, S.; Drost, T.; Dirksen, Wessel; Menendez, M.; Nguyen, H.; Phelps, M.; Knopp, M.V.; Rosol, T. J. “A versatile canine prostate cancer model that expresses human receptors” James CCC Translational Therapeutics retreat. Oct 31, 2015, Columbus, Ohio.**

23. **Tweedle, M.F., C Beuttner, A Wallace, M Williams, K Kumar, JE Goldberger**  
**Gd chelates that self-assemble in a low pH biological microenvironment. Contrast**  
**Media Research 2015, Berlin , Germany**

24. **John Phay, Steven Justiano, Haiming Ding, Kara Rossfeld, Shankaran**  
**Kothandaraman, Chadwick Wright, Matthew Ringel, Michael Tweedle. A novel**  
**imaging agent for medullary thyroid carcinoma in an orthotopic mouse model.**

### External lectures

1. **2015 Case Comprehensive Cancer Center. “A versatile Canine Prostate Cancer Model” at Cancer Imaging Mini-Retreat, Friday October 16, 2015.**
2. Case Comprehensive Cancer Center. “Targeting Peptide Diagnostic Pharmaceuticals with Commercial Intent. November, 2013.
3. **Media Contrast Research 2015, Berlin, Germany “Gd chelates that self-assemble in a low pH biological microenvironment. Contrast”**

### Internal Lectures

1. **Engineering 4891. Creativity and Innovation. “Entrepreneurial Attitudes.” November, 2015.**
2. **James CCC Grand Rounds. “Pan-Cancer Targeting MRI Contrast Agents that Self-assemble in Malignant Tumors.” May, 2015.**
3. **Residents Seminar. “History and Pharmacology of X ray and MRI Contrast Agents.” July, 2015.**
4. **Radiology Safety Committee talk. “GBCA and Residual Gd.” September, 2015.**
5. **Drug Development Institute. “Gd PA molecules as pan cancer imaging agents.” December 2015.**

6. **Residenmts Seminar**

### Educational Video

1. "ProHance, new Imaging Pharmaceutical for MRI." Bristol-Myers Squibb training VHS tape, 1989 (est.)
2. Prescription New Jersey, hosted by Sara Lee Kessler. "New Jersey Network Television documentary" Last aired July 28, 2005
3. 2011 Stefanie's Champions Awards Ceremony – "Continue to Fight"  
<http://www.youtube.com/watch?v=NLDu94y3g6M>
4. 2010 Stefanie's Champions Awards Ceremony – Opening Video  
<http://www.youtube.com/watch?v=-XoN9GA-mGU&feature=plcp>
5. **Radiological Society of North America. Interview and discussion. June 2015. Release.**

**Recent Press Reports (2009 – 2015 only) (Two new in 2013/14)**

1. Columbus Post dispatch Article on Stefanie Spielman  
[http://www.dispatch.com/live/content/local\\_news/stories/2009/11/24/Spielman\\_Funds.ART\\_ART\\_11-24-09\\_A1\\_FGFPJEU.html](http://www.dispatch.com/live/content/local_news/stories/2009/11/24/Spielman_Funds.ART_ART_11-24-09_A1_FGFPJEU.html)
2. WBNS-10/TV (Komen Race for the Cure special)  
As part of a 30-minute "behind-the-scenes" look at the upcoming Race for the Cure special, OSUCCC cancer researcher **Dr. Michael Tweedle** discusses his on-going research to develop a drug that would allow surgeons to know in real-time during surgery if they had removed all of the cancerous cells. "The drug would be like a bloodhound seeking out cancerous cells," said Tweedle, "and would glow when illuminated by a special light, thus allowing surgeons to see if they had clear margins before finishing the surgery."  
WBNS: <http://bit.ly/ITCXAT>
3. SNM Smart Brief with Jeff Kurate and Michael F. Tweedle  
Fall 2013 Re; Tauopathic Imaging Agents  
"Researchers create "designer" tracer for earlier diagnosis and better treatments of Alzheimer's" "Unlike beta amyloid, tau appears in specific [brain](#) regions in Alzheimer's," said Tweedle. "With a better view of how tau is distinct from amyloid, we'll be able to create a much more accurate view of disease staging, and do a much better job getting the right therapeutics into the right populations at the right time."  
<http://www.news-medical.net/news/20130426/Researchers-create-designer-tracer-for-earlier-diagnosis-and-better-treatments-of-Alzheimers.aspx>
4. **Internal CCTS Interview Online with Chad Wright and Michael Tweedle, Spring 2014 Re: Bremer Grant** "<sup>123</sup>Iodine-labeled HN1-800dye agents for dual-modality intraoperative surgical guidance during resection of head & neck squamous cell carcinoma." "Before a surgeon does any cutting, they will do imaging, like PET/CT, MRI or CT scans, to try to locate the cancer," Tweedle said. Wright and Tweedle's efforts have been strengthened by an outstanding team of multidisciplinary collaborators, including Nathan Hall, MD, PhD, Quintin Pan, PhD,

**Ted Teknos, MD, and Michael Knopp, MD, PhD. Their novel, real-time imaging detection and treatment paradigm is critical to improving and personalizing cancer detection and care at Ohio State.** <https://ccts.osu.edu/node/4171>

**Optical Surgical Navigation workshop at the World Molecular Imaging Congress, co- organized** <http://www.wmis.org/optical-surgical-navigation-workshop-at-the-wmic-2014/>

**Synopsis.** <http://www.wmis.org/wp-content/uploads/2014/10/WMIC-2014-OSN-Workshop-RIs-Final.pdf>