Harvard Medical School/Harvard School of Dental Medicine

| Date Prepared: | February, 2018 |
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| Name: | Irene Emily Kochevar |
| Office Address: | Wellman Center for Photomedicine Department of Dermatology Massachusetts General Hospital, Thier 212 Harvard Medical School Boston, MA 02114 |
| Work Phone: | 617-726-8828 |
| Work Email: | kochevar@helix.mgh.harvard.edu |
| Work FAX: | 617-726-3192 |
| | |

Education

| 1965 | BS | Biochemistry | Michigan State University |
|------|-----|--------------|---------------------------|
| 1967 | MS | Biochemistry | Michigan State University |
| 1970 | PhD | Chemistry | Michigan State University |

Postdoctoral Training

| 1970-1971 | NIH Postdoctoral Fellow | Department of Chemistry | New York University |
|-----------|----------------------------|-------------------------|---------------------|
| 1975-1977 | NIH Postdoctoral Fellow | Department of Chemistry | Columbia University |

Faculty Academic Appointments

| Research Associate | Department of Chemistry | Columbia University |
|--|--|---|
| Research Associate | Department of Biochemistry | Columbia University |
| Assistant Professor of Dermatological | Department of Dermatology | Columbia University |
| Dringing LAggesists in | Department of Demostele av | Hawand Madical Sabaal |
| Dermatology | Department of Dermatology | Harvard Medical School |
| Visiting Professor | Department of Chemistry | University of Connecticut |
| Associate Professor | Department of Dermatology | Harvard Medical School |
| Associate Professor | Division of Health Sciences & | Massachusetts Institute of |
| | Technology | Technology |
| Professor | Department of Dermatology | Harvard Medical School |
| Associated Faculty | Harvard-MIT Health Sciences Technology Program | Massachusetts Institute of Technology |
| | Research Associate Research Associate Assistant Professor of Dermatological Biochemistry Principal Associate in Dermatology Visiting Professor Associate Professor Associate Professor Professor Associated Faculty | Research AssociateDepartment of ChemistryResearch AssociateDepartment of BiochemistryAssistant Professor ofDepartment of DermatologyDermatologicalDepartment of DermatologyBiochemistryDepartment of DermatologyPrincipal Associate inDepartment of DermatologyDermatologyDepartment of DermatologyVisiting ProfessorDepartment of ChemistryAssociate ProfessorDepartment of DermatologyAssociate ProfessorDepartment of DermatologyProfessorDepartment of DermatologyProfessorDepartment of DermatologyAssociate ProfessorDepartment of DermatologyAssociated FacultyHarvard-MIT Health Sciences Technology Program |

Appointments at Hospitals/Affiliated Institutions

| 1981-1986 | Associate Biochemist | Massachusetts General Hospital |
|-----------|----------------------|--------------------------------|
| 1986- | Biochemist | Massachusetts General Hospital |

Other Professional Positions

1971-1973 Research Chemist Union Carbide Corporation, Bound Brook, NJ

Major Administrative Leadership Positions

| Local | | |
|-----------|--------------------|--------------------------------------|
| 1965-1970 | Research Assistant | Departments of Biochemistry and |
| | | Chemistry, Michigan State University |

National and International

| 1988-1992 | Association Internationale de Photobiologie | Vice President |
|-----------|---|--|
| 1980-1983 | American Society for Photobiology | Council member |
| 1985-1987 | American Society for Photobiology | President-Elect, President, Past President |
| 1998-2002 | Inter-American Photochemical Society | Board of Directors |

Committee Service

Local

1983-1987Subcommittee on Research SafetyMassachusetts General Hospital1986-1992,Subcommittee on Review of ResearchMassachusetts General Hospital1997-2000ProposalsMassachusetts General Hospital1986-1987,Committee on ResearchMassachusetts General Hospital1990-1993Massachusetts General Hospital

National and International

| 1979-1983 | Photobiology Task Force | American Academy of Dermatology |
|-------------|----------------------------|---|
| 1979-1981 | Committee on Photobiology | National Research Council |
| 1980-1983 | Council | American Society for Photobiology |
| 1985 | National Meeting Organizer | American Society for Photobiology |
| 1986-1991 | Advisory Board | Center for Fast Kinetics Research, |
| | | University of Texas at Austin |
| 1992-1996 | Scientific Advisory Board | Center for Advanced Research in |
| | | Photobiology |
| 1991, 1995, | Scientific Advisory Board | NIEHS Laboratory of Molecular Biophysics |
| 2004 | | |
| | | |
| 2007-2013 | Scientific Advisory Board | Chemistry, Chemical Biology and |
| | | Biomedical Engineering, Stevens Institute |
| | | of Technology |

Professional Societies

| American Chemical Society | Member |
|--|--------|
| American Society for Photobiology | Member |
| Inter-American Photochemical Society | Member |
| Society for Investigative Dermatology | Member |
| New York Academy of Sciences | Member |
| American Association for the Advancement | Member |
| of Science | |
| European Photochemical Association | Member |
| European Society for Photobiology | Member |
| Society for Free Radicals in Biology and | Member |
| Medicine | |

Grant Review Activities

1994, 1995NIH MBRC Evaluation committeesNIH1997 - 2001Regular memberNIH General Medicine A-1 Study SectionReviewer for many NIH special review panels and multiple European and Canadian granting agencies;list not recorded.

Editorial Activities

Reviewer for many journals covering topics in organic chemistry, photochemistry, photobiology, dermatology, photomedicine, oxidative stress, cell biology, laser medicine.

Other Editorial Roles

| 1983-1993 | Associate Editor | Photochemistry and Photobiology |
|-----------|------------------|------------------------------------|
| 1994-1998 | Editor-in-Chief | Photochemistry and Photobiology |
| 1990-1999 | Associate Editor | Photodermatology, Photoimmunology, |
| | | Photomedicine |

Honors and Prizes

| 1967 | Dow Summer Fellow | Department of Chemistry, Michigan State University |
|-----------|---------------------------------------|---|
| 1967-1968 | Du Pont Fellow | Department of Chemistry, Michigan State University |
| 1968 | Outstanding Woman Graduate Student | Michigan State University |
| 1969-1970 | Lubrizol Fellow | Department of Chemistry, Michigan State University |
| 1995 | Fellow | American Association for the Advancement of Science |
| 1997 | Photon Award | American Society for Photobiology |

Report of Local Teaching and Training

Teaching of Students in Courses

| 1978-1980 | Biochemistry | Columbia University |
|---|---|---|
| | | Organized and taught biochemistry seminar |
| | | for medical students |
| 1983, 1985, | Harvard University course on Photobiology | Lecturer, Harvard University |
| 1989, 1991 | (Biology 289) | |
| 1988-1990 | HST 569, Photomedicine | Massachusetts Institute of Technology, |
| | | Course Director |
| 1992 | HST 569, Photomedicine | Massachusetts Institute of Technology, |
| | | Course Director |
| | | |
| Formal Teaching of Residents, Clinical Fellows and Research Fellows (post-docs) | | |

| 1977-1981 | Photobiology Seminar | Columbia University Organized and lectured in course for dermatology residents and fellows |
|-----------|---|--|
| 1983 | Photobiology Course for Dermatology residents | Lecturer, Massachusetts General Hospital |
| 1985-1988 | Wellman tutorials for residents and fellows | Massachusetts General Hospital series of six sessions each year in photochemistry and photobiology |
| 1989-1990 | Photobiology Workshops and Lecture Series | Massachusetts General Hospital, Co- Coordinator |

Laboratory and Other Research Supervisory and Training Responsibilities

Formally Supervised Postdoctoral Trainees (records not kept for undergraduate trainees)

| Training Period | Trainee Name | Degree(s) | Research Proiect Title | Current Position of Past Trainees |
|--------------------|---------------------|-----------|-----------------------------|--------------------------------------|
| 1979-1980 | Chung, Fung Lung | PhD | Chlorpromazine photo- | Prof. Oncology, |
| | | | toxicity | Georgetown Univ. |
| 1978-1980 | Gasparro, Frank | PhD | Protryptyline phototoxicity | Chem. teacher, Hall |
| | | | | Hamden School, CT |
| 1980-1982 | Yoon, MinJoun | PhD | UV-induced membrane | Prof. Chem., Seoul Natl. |
| | | | damage | Univ., Korea |
| 1983-1984 | Gendimenico, Gerard | PhD | Mast cells in drug photo- | Sr. Proj. Mgr., Johnson & |
| | | | toxicity | Johnson |
| 1983 | Hasan, Tayyaba | PhD | Tetracycline drug photo- | Prof. of Dermatology, |
| | | | toxicity | HMS; Wellman |
| | | | | Photomed., MGH |
| 1985-1987 | Van Camp, John | PhD | Rose Bengal photosen- | Senior Research Leader, |
| | | | sitized DNA cleavage | Perkin Elmer, |
| | | | mechanisms | Mountainview CA |
| 1985-1986 | Western, Andre | PhD | Amiodarone drug photo- | CRA, Quintiles; Study |
| | | | toxicity | coord., Nycomed |
| | | | | Imaging, Norway |

| 1987-1988 | Valdes-Aguilera, | PhD | Photosenstized mitochon- | Sr. Development |
|------------|----------------------|----------|------------------------------|-----------------------------|
| | Oscar | | drial damage | Chemist, Cytec Surface |
| | | | C C | Industries, GA |
| 1987-1989 | Hurley, John | PhD | Laser flash photolysis of | Senior Research Assoc. |
| | | | drugs | University of Arizona |
| 1988-1990 | Dunn, David | PhD | Dye photosensitized DNA | Dir., HTS Technologies, |
| | , | | cleavage | Pharmacopeia, Inc. |
| 1988-1989 | Allen, Mary Tedd | PhD | Rose Bengal membrane | VP Manufacturing, |
| | | | photosensitization | nanoString Technology, |
| | | | 1 | Seattle WA |
| 1989-1992 | Redmond, Robert W. | PhD | Multiphoton | Assoc Prof, Wellman |
| | | | photosensitization | Center, MGH/HMS |
| 1989-1991 | Chaudry, Nina | MD | Photovasorelaxation | Asst. Prof. Cardiology, |
| | | | | Columbia Univ. Coll. of |
| | | | | Physicians and Surgeons, |
| | | | | NY, NY |
| 1988-1990 | Lilge, Lothar | PhD | Applications of light + | Assoc. Prof./Sr. Scientist, |
| | | | lasers in medical pre- | Ontario Cancer |
| | | | clinical + clinical research | Inst./Princess Margaret |
| | | | | Hosp., Toronto, Can. |
| 1991-1994 | Garcia, Carmello | PhD | Chlorpromazine photo- | Prof. Chem., Univ. |
| | | | physics + photochemistry | Puerto Rico |
| 1992-1995 | Nguyen, Bach-Cuc | PhD | Mechanisms for skin | Research Assoc., CBRC, |
| | | | pigmentation induced by | MGH |
| | | | dihydroyacetone | |
| 1993-1994 | Lambert, Christopher | PhD | Intensity dependent photo- | Assoc. Research Prof., |
| | | | chemistry and photo- | Bioen-gineering Inst., |
| | | | biology of rose Bengal | Worcester Polytechnic |
| 1000 1007 | | | | Inst., MA |
| 1993-1995 | Martinez, Lydia | PhD | Two-photon DNA photo- | Visiting Prof. Chem., |
| 1004 100 6 | | | chemistry | Union Coll., NY |
| 1994-1996 | Gonzalez, Salvador | MD | Involvement of mast cells | Assoc. Prot. of Med., |
| | | | in chronic UVB damage to | Univ. Alcala, Madrid, |
| 1004 1007 | | DI D | SKIN | Spain D. C. LL : C |
| 1994-1995 | Tedesco, Antonio C. | PnD | Photosensitized plasma | Prof. Chem., Univ. of |
| 1005 1007 | Dhiling Maana | DLD | Effects of change | Sao Paolo, Brazil |
| 1995-1997 | Philips, Neena | PnD | Effects of chronic UVB on | Assoc. Prol. of Biol. |
| | | | dermai horobiasts | Scis., Fairleign |
| 1006 | Ibbotoon Colly | MD | Error radicals and skin | Sr. Lasturer Ninewalla |
| 1990 | 1000tson, Sany | | aging | Hosp & Mod School |
| | | | agnig | Dundee Scotland |
| 1008 2002 | Zhuang Shougang | որ որ որ | Singlet ovygen induged | Assoc Prof Brown |
| 1770-2003 | Zinuang, Shougang | | signal transduction path | Liniv Med School |
| | | | ways | |
| 2000-2002 | Chan Barbara P | PhD | Photochemical tissue | Assoc Prof |
| 2000-2002 | Chain, Darbara I. | | bonding for skin grafts and | Bioengineering Univ of |
| 1 | | 1 | 1 oonung ior skin grans allu | Divengineering, Ulliv. Ul |

| | | | tendon repair | Hong Kong |
|-----------|----------------------|---------|--------------------------------|---------------------------|
| 2000-2002 | Proano, Cinthia | MD | Photochemical tissue bond- | Faculty, Essex |
| | | | ing of cornea in vivo | Community Coll. |
| 2001-2002 | Mitsomoto, Atushi | PhD | Role of reactive oxygen | Assoc. Prof., Josai Intl. |
| | | | species in stimulation of | Univ., Tokyo, Japan |
| | | | elastin produced by dermal | |
| | | | fibroblasts | |
| | | | | |
| 2001-2003 | Kossodo, Sylvie | PhD | Effects of UV-induced | Research Group Leader, |
| | | | cytokines on extracellular | VisEn Med. Co., |
| | | | matrix proteins + proteases | Bedford, MA |
| 2001-2005 | Choi, Won-Seon | PhD | Role of TGF- β in skin | Patent Lawyer, |
| | | | photoaging | Washington DC |
| 2002-2003 | Wong, Wen-Rou | MD | Involvement of UV- | Clin. practice, Chang- |
| | | | induced cytokines in | Gung Med. Coll., Taiwan |
| | | | photoaging | _ |
| 2002-2004 | Carle, A. Bjoern | PhD | Selective UVA-induced | Sr. Applications |
| | - | | processes at the plasma | Scientist, Artel, Inc. |
| | | | membrane using evanes- | |
| | | | cent wave radiation | |
| 2002-2005 | Kamegaya, Yoko | MD | Evaluation of photochemi- | Dept. of Dermatol., Keio |
| | | | cal tissue bonding for | Univ. Hosp., Japan |
| | | | closure of skin incisions + | |
| | | | excisions | |
| 2003-2005 | Wang, Hongjun | PhD | UVB-induced production | Assoc. Prof. Biomed. |
| | | | of TGF- β by human kera- | Engineering, Stevens |
| | | | tinocytes | Inst. of Technology, |
| | | | | Hoboken, NJ |
| 2003-2006 | Valencia, Antonio | PhD | Mechanisms of apoptotic | Group Leader, |
| | | | cell death by free radicals | Biogen/Idec, Cambridge |
| | | | | MA |
| 2005-2006 | Dowdall, Jayme Rose | MD | Photochemical repair of | Asst. Prof, |
| | | | vocal fold microflap | Otolaryngology, MEEI |
| | | | defects | |
| 2006-2008 | Yamaura, Maki | MD | Role of superoxide genera- | Dept. Dermatol., Shinshu |
| | | | ting NADPH oxidase 4 in | Univ. Hosp., School of |
| | | | melanotic cell proliferation | Med Japan |
| 2006-2010 | Yao, Min | MD, PhD | Technology for sutureless | Prof. Burn & Plastic |
| | | | closure of surgical | Surgery, Shanghai Jio |
| | | | incisions + wounds | Tong University Medical |
| | | | | School, China |
| 2007 | Chung, Connie | MD, PhD | Biophysics + protein | Clin. Instr. Dermatol., |
| | | | structure | HMS; Dermatologist, |
| | | | | MGH |
| 2008-2009 | Wang, Ying | MD | Light-activated treatment | Instructor, Wellman |
| | | | for ocular surface damage | Center, MGH |
| 2008-2009 | Venditti, Elisabetta | PhD | UVA and nitroxides as | Research Assistant at |

| | | | inhibitors of oxidative | Istituto Europeo di |
|-----------|--------------------|---------|-----------------------------|--------------------------|
| | | | damage to skin collagen | Oncologia (IEO) |
| 2008-2011 | Datta, Neha | MD | Photochemical | Resident, Dept Surgery, |
| | | | technologies for surgical | UCLA |
| | | | applications | |
| 2009-2010 | Tang, Yijin | PhD | Collagen photocrosslinking | Research Fellow, New |
| | | | mechanisms | York State Department |
| | | | | of Health |
| 2010-2011 | Gu, Chuan | MD | Photorepair of limbal stem | Dept. of Plastic & Burn |
| | | | cell deficiency | Surg., No. 3 People's |
| | | | | Hosp., Shanghai Jiao |
| | | | | Tong University, |
| | | | | Shanghai China |
| 2010-2011 | Bajwa, Amrita Kaur | MA | Photobonding to repair | Grad. Student, Dept. |
| | | | penetrating eye wounds | Plant + Microbial Biol., |
| | | | | Univ. CA, Berkeley |
| 2010-2011 | Yang, Penggao | MD | Effects of photochemical | Dept. of Plastic & Burn |
| | | | tissue bonding on one-layer | Surg., No. 3 People's |
| | | | skin incision closure; | Hosp., Shanghai Jiao |
| | | | | Tong University, |
| | | | | Shanghai China |
| 2012 | Tjiu, Jengwei | MD, PhD | Photosensitization of | Asst. Prof., Derm. |
| | | | keratinocytes | National Taiwan Univ. |
| 2012-2015 | Zhu, Hong | MD | Keratoconus treatment | Ophthalmology, People's |
| | | | | Hospital #1, Shanghai, |
| | | | | China |
| 2012-2015 | Xu, Nan | MD | Pigmented skin wound | Shanghai East Hospital, |
| | | | sealing | Tongji University School |
| | | | | of Medicine, Shanghai, |
| | | | | China |
| | | | | |
| 2016- | Seiler, Theo G | MD | Interfacial cornea | |
| present | | | photobonding | |
| 2017-2018 | Wertheimer, C. | MD | Crosslinking mechanisms | |

Formal Teaching of Peers (e.g., CME and other continuing education courses)

| 1979-1982 | Photobiology course, taught basic photobiology | American Academy of |
|-----------|--|--------------------------------|
| | | Dermatology Meeting, Lecturer |
| 1980 | Photobiology course | Columbia University |
| | | Organized and lectured in six- |
| | | hour course |
| 1982-1984 | Photobiology Workshops and Lecture Series, | Massachusetts General Hospital |
| | Coordinator | |

Local Invited Presentations

Record not maintained of presentations.

<u>Report of Regional, National and International Invited Teaching and</u> <u><u>Presentations</u></u>

Invited Presentations and Courses

Record not maintained of presentations.

Report of Technological and Other Scientific Innovations

| Photochemical tissue bonding | US 8092490 | January 10, 2012 |
|---------------------------------|---------------|-------------------|
| Photochemical tissue | US 8215314 | July 10, 2012 |
| Photochemical tissue | US 7331350 | February 19, 2008 |
| bonding | | - |
| Photochemical tissue | US 7073510 | July 11, 2006 |
| bonding | | |
| Photochemical tissue | AU 2001249984 | |
| bonding | | |
| Photochemical tissue | JP 5101778 | |
| bonding | | |
| Photochemical tissue | EP 1272119 | |
| bonding | | |
| Photochemically | EP 3067015 | |
| induced engagement of | | |
| intraocular implants | | |

Report of Scholarship

Publications

Peer reviewed publications in print or other media

- 1. Speck JC Jr, Rynbrandt DJ, Kochevar IE. Neighboring group participation in acetal hydrolysis. J Am Chem Soc. 1965; 87:4979-4980.
- 2. Wagner PJ, Kochevar IE. How efficient is diffusion controlled triplet energy transfer? J Am Chem Soc. 1968; 90:2232-2238.
- 3. Kochevar IE. A kinetic study of the quenching of triplet butyrophenone by mono-olefins. Ph.D. Dissertation. Michigan State University, 1970.
- 4. Kochevar IE, Wagner PJ. Triplet ketone-olefin interactions: Energy transfer, charge transfer or radical addition? J Am Chem Soc. 1970; 92:5742-5743.

- 5. Kochevar IE, Wagner PJ. Quenching of triplet phenyl ketones by olefins. J Am Chem Soc. 1972; 94:3859-3865.
- Wagner PJ, Kochevar IE, Kempainin AE. Type II photoprocesses of phenyl ketones. Procedure for determining meaningful quantum yields and triplet lifetimes. J Am Chem Soc. 1972; 94:7489-7495.
- 7. Wamser CC, Medary RT, Kochevar IE, Turro NJ, Chang PL. Steric effects on singlet and triplet energy transfer to azo compounds. J Am Chem Soc. 1975; 97:4864-4869.
- 8. Kochevar IE, Harber LC. Photoreactions of 3,3',4',5-tetrachloroalicylanilide with proteins. J Invest Dermatol. 1977; 68:151-156.
- 9. Turro NJ, Kochevar IE, Noguchi Y, Chow MF. Electronic excitation transfer in polymers III. Singlet-singlet, triplet-singlet and triplet-triplet energy transfers. Evidence for triplet migration among pendant phenyl groups in polystyrene. J Am Chem Soc. 1978; 100:3170-3177.
- 10. Kochevar IE. Photoallergic responses to chemicals. Photochem Photobiol. 1979; 30:437-442.
- 11. Kochevar IE, Lamola AA. Chlorpromazine and protriptyline phototoxicity: photosensitized, oxygen independent red cell hemolysis. Photochem Photobiol. 1979; 29:791-796.
- 12. Kochevar IE, Zalar GL, Einbinder J, Harber LC. Assay of contact photosensitivity to musk ambrette in guinea pigs. J Invest Dermatol. 1979; 73:143-146.
- 13. Sutherland BM, Harber LC, Kochevar IE. Pyrimidine dimer formation and repair in human skin. Cancer Res. 1980; 40:3181-3185.
- 14. Kochevar IE. Toxicity due to photochemical products of protriptyline. Toxicol Appl Pharmacol. 1980; 54:258-264.
- 15. Giovinazzo VJ, Harber LC, Armstrong RB, Kochevar IE. Photoallergic contact dermatitis to musk ambrette: Clinical report of two patients with persistent light reactor patterns. J Am Acad Dermatol. 1980; 3:384-393.
- Giovinazzo VJ, Ichikawa H, Kochevar IE, Armstrong RB, Harber LC. Photoallergic contact dermatitis to musk ambrette: Action spectra in guinea pigs and man. Photochem Photobiol. 1981; 33:773-777.
- 17. Lamola AA, Landon D, Kochevar IE, Harber LC. An instrument for action spectrum studies in dermatology. Photochem Photobiol. 1982; 35:285-290.
- 18. Gasparro FP, Kochevar IE. Investigation of protriptyline photoproducts which cause cell membrane disruption. Photochem Photobiol. 1982; 35:351-358.
- 19. Kochevar IE, Armstrong RB, Einbinder J, Walther RR, Harber LC. Coal tar phototoxicity: active compounds and action spectra. Photochem Photobiol. 1982; 36:65-69.

- 20. Kochevar IE, Hom J. Photoproducts of chlorpromazine which cause red blood cell lysis. Photochem Photobiol. 1983; 37:163-168.
- 21. Kochevar IE, Yoon M. Photosensitization of methyl linoleate oxidation by tryptophan in peptides. Photochem Photobiol. 1983; 37:279-286.
- 22. Hasan T, Kochevar IE, McAuliffe DJ, Cooperman BS, Abdulah D. Mechanism of tetracycline phototoxicity. J Invest Dermatol. 1984; 83:179-183.
- 23. Kochevar IE, Wujek Hoover K, Gawienowski M. Benoxaprofen photosensitization of cell membrane disruption. J Invest Dermatol. 1984; 82:214-218.
- 24. Kochevar IE, Chung F-L, Jeffrey AM. Photoaddition of chlorpromazine to DNA. Chem Biol Interact. 1984; 51:273-284.
- 25. Gendimenico GJ, Kochevar IE. Degranulation of mast cells and inhibition of the response to secretory agents by phototoxic compounds and ultraviolet radiation. Toxicol Appl Pharmacol. 1984; 76:374-382.
- 26. Hasan T, Kochevar IE, Abdulah D. Amiodarone phototoxicity to human erythrocytes and lymphocytes. Photochem Photobiol. 1984; 40:715-719.
- 27. Kochevar IE. Influence of prior complex formation on the photoaddition of chlorpromazine to calf thymus DNA. J Photochem. 1985; 28:195-201.
- 28. Gange RW, Blackett AD, Matzinger E, Sutherland BM, Kochevar IE. Comparative protection efficacy of UVA and UVB induced tans against erythema and pyrimidine dimer induction by UVB in human skin. J Invest Dermatol. 1985; 85:362-364.
- 29. Kochevar IE, Morison WL, Lamm JL, McAuliffe DJ, Western A, Hood AF. Possible mechanism of piroxicam-induced photosensitivity. Arch Dermatol 122:1283-1287, 1986.
- 30. Ciulla TA, Epling GA, Kochevar IE. Photoaddition of chlorpromazine to guanosine-5'monophosphate. Photochem Photobiol. 1986; 43:607-613.
- 31. Valdes-Aguilera O, Cincotta L, Foley J, Kochevar IE. Photobleaching of a cyanine dye in solution and in membranes. Photochem Photobiol. 1987; 45:337-344.
- 32. Green H, Boll J, Parrish JA, Kochevar IE, Oseroff AR. The cytotoxicity and mutagenicity of low intensity 248 and 193 nm excimer laser radiation in mammalian cells. Cancer Res. 1987; 47:410-413.
- 33. Green H, Margolis RJ, Boll J, Kochevar IE, Parrish JA, Oseroff AR. Unscheduled DNA synthesis in human skin after in vitro ultraviolet-excimer laser ablation. J Invest Dermatol. 1987; 89:201-204.

- 34. Western A, Van Camp JR, Bensasson R, Land EJ, Kochevar IE. Involvement of singlet oxygen in the phototoxicity mechanism for a metabolite of piroxicam. Photochem Photobiol. 1987; 46:469-475.
- 35. Valdes-Aguilera O, Ara G, Kochevar IE. Phototoxicity mechanism of a kryptocyanine dye in red blood cell membranes and isolated mitochondria. Cancer Res. 1988; 48:6794-6798.
- 36. Ciulla TA, Van Camp JR, Rosenfeld E, Kochevar IE. Photosensitization of single-strand breaks in pBR322 DNA by rose bengal. Photochem Photobiol. 1989; 49:293-299.
- Fluhler EN, Hurley JK, Kochevar IE. Laser intensity and wavelength dependence of rose bengal photosensitized inhibition of red cell acetylcholinesterase. Biochim Biophys Acta 1989; 990:269-275.
- 38. Gallo RL, Kochevar IE, Granstein RD. Ultraviolet radiation induces a change in cell membrane potential in vitro: A possible signal for ultraviolet radiation induced alteration in cell activity. Photochem Photobiol. 1989;49:655-662.
- 39. Gendimenico GJ, Kochevar IE. A further characterization of acridine-photosensitized inhibition of mast cell degranulation. Photoderm Photoimmunol Photomed. 1990;7:51-5.
- 40. Kochevar IE, Walsh AA, Held KD, Gallo RL, Mirro J. Mechanism for 193-nm laser radiationinduced effects on mammalian cells. Radiation Res. 1990;122:142-148.
- 41. Kochevar IE, Buckley LA. Photochemistry of DNA using 193 nm excimer laser radiation. Photochem Photobiol. 1990;51:527-532.
- 42. Kochevar IE. UV-induced protein alterations and lipid oxidation in erythrocyte membranes. Photochem Photobiol. 1990;52:795-800.
- 43. Hefetz Y, Dunn DA, Deutsch TF, Buckley L, Hillenkamp F, Kochevar IE. Laser photochemistry of DNA: Two-photon absorption and optical breakdown using high intensity, 532-nm radiation. J Am Chem Soc. 1990;112:8528-8532.
- 44. Andley UP, Walsh A, Kochevar IE, Reddan JR. Effect of UVB radiation on protein synthesis in cultured lens epithelial cells. Current Eye Res. 1990;9:1099-1106.
- 45. Dunn DA, Lin VH, Kochevar IE. The role of ground state complexation in the electron transfer quenching of methylene blue fluorescence by purine nucleotides. Photochem Photobiol. 1991; 53:47-56.
- 46. Kochevar IE, Walsh AA, Green HA, Sherwood M, Shih AG, Sutherland BM. DNA damage induced by 193-nm radiation in mammalian cells. Cancer Res. 1991;288-293.
- 47. Allen MT, Lynch M, Lagos A, Redmond RW, Kochevar IE. A wavelength dependent mechanism for rose bengal-sensitized photoinhibition of red cell acetylcholinesterase. Biochim Biophys Acta 1991; 1075:42-49.

- 48. Dunn DA, Lin VH, Kochevar IE. Base selective oxidation and cleavage of DNA by photochemical cosensitized electron transfer. Biochemistry 1992;31:11620-11625.
- 49. Kochevar IE, Moran M, Lyon N, Flotte F, Siebert E, Gange RW. Effects of systemic indomethacin, meclizine and BW755C on chronic UVB-induced effects in hairless mouse skin. J Invest Dermatol. 1993;100:186-193.
- 50. Lilge L, Flotte TJ, Kochevar IE, Jacques SL, Hillenkamp F. Photoactivable fluorophores for the measurement of fluence in turbid media. Photochem Photobiol 1993;58:37-44.
- 51. Gut IG, Hillenkamp F, Hefetz Y, Kochevar IE. Two-photon absorption cross-sections of guanosine 5'-monophosphate and uridine 5'-monophosphate at 532 nm. J Phys Chem. 1993; 97:5171-5176.
- 52. Gut IG, Farmer R, Huang RC, Kochevar IE. Upper excited state photochemistry of DNA. Photochem Photobiol 1993;58:313-317.
- 53. Chaudhry H, Lynch M, Schomacker K, Birngruber R, Gregory K, Kochevar I. Relaxation of vascular smooth muscle induced by low power laser radiation. Photochem Photobiol 1993; 58:661-669.
- 54. Smith, G, McGimpsey WG, Lynch MC, Kochevar IE, Redmond, RW. An efficient oxygen independent two-photon photosensitization mechanism. Photochem Photobiol 1994;59:135-139.
- 55. Andley UP, Lewis RM, Reddan JR, Kochevar IE. Action spectrum for cytotoxicity in the UVAand UVB-wavelength region in cultured lens epithelial cells. Invest Ophthal Vis Sci 1994; 35:367-373.
- 56. Kochevar IE, Moran M, Granstein RD. Experimental photoaging in C3H/HeN, C3H/HeJ, and Balb/c mice. Comparison of changes in extracellular matrix components and mast cell numbers. J Invest Dermatol 1994;103:797-800.
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Thesis

A kinetic study of the quenching of triplet ketenes by mono-olefins. Michigan State University, 1970.

Abstracts, Poster Presentations and Exhibits Presented at Professional Meetings

No record kept of abstracts and posters.

Narrative Report (limit to 500 words)

My laboratory has focused on understanding the normal and pathological effects of light (UV/visible) on human tissue (mainly skin) and using this knowledge to develop light-initiated therapies. Educated in physical organic chemistry, photochemistry and biochemistry, I began my independent research career more than 35 years ago. Starting with studies of the photochemical mechanisms underlying drug phototoxicity responses, the research evolved toward phototherapies. Concurrently we studied the effects of UV radiation on cells with special emphasis on cell signaling initiated at the plasma membrane and processes leading to chronic UV-induced skin damage. For the last 10 years, in collaboration with Robert Redmond PhD at the Wellman Center, we developed a light-activated technology for tissue repair based on photo-crosslinking tissue proteins. Studies carried out ex vivo and in vivo in tissues have demonstrated its utility for a wide range of tissue repair and examined the molecular mechanism.

Drug phototoxicity /therapeutic photosensitization. Both of these phenomena, although one is deleterious and the other beneficial, involve formation and reactions of singlet oxygen. One focus has been the formation of singlet oxygen in and near the cell plasma membrane, for example, by selective delivery of light to this region using evanescent waves. We established signaling pathways by which singlet oxygen stimulates and/or damages cells. Examples include: 1) Demonstrated differences in cell death mechanisms initiated by singlet oxygen versus hydrogen peroxide. 2) Established that singlet oxygen down-regulates EGFR signaling by activating caspase-3 and protein phosphatases. 3) Determined that protein kinase C inhibits singlet oxygen-induced apoptosis by decreasing caspase-8 activation.

<u>UV radiation-initiated responses.</u> In addition to the mutagenic effects of UV that are initiated by DNA damage, reactive oxygen species (ROS) are produced in skin exposed to solar UV. These ROS contribute to acute effects (inflammation) and chronic damage (photoaging and photocarcinogenesis). Our goal was to reduce the damaging effects of ROS on skin. Two examples: 1) While studying the mechanisms for the UVA photosensitivity in Smith-Lemli-Optiz syndrome, we identified Nox1 as a source of UVA-initiated ROS formation in keratinocytes thus providing a therapeutic target. 2) In investigations of solar elastosis, a prominent sign of chronic skin photodamage, we established that ROS mediate TGF- β initiated elastin production.

Light-activated tissue repair. This project was initiated in response to the need for a rapid, water-tight, sutureless method for closing skin wounds and other damaged tissue requiring adhesion between tissue surfaces. We demonstrated that light-activated linking of proteins between the tissue surfaces addressed this need. In Photochemical Tissue Bonding (PTB) a photoactive dye is applied to tissue surfaces and irradiated with visible light to covalently crosslink proteins without toxicity or inflammation. PTB was superior to sutures for closure of skin wounds (including a 30-patient study) and for reconnecting peripheral nerves, blood vessels and tendons in animal studies. In cornea, PTB rapidly sealed penetrating wounds and attached amniotic membrane transplants and limbal stem cells. More recently we have used light-activated protein crosslinking within, rather than between, tissues to strengthen the tissue or change its properties. An example is photo-crosslinking collagen in cornea as a treatment for keratoconus. Light-

activated tissue repair remains an active area of research including identifying mechanisms in order to enhance the efficiency of PTB.