

2007 Meeting of the
Society for
Whole-Body Autoradiography

March 18-21, 2007

**Mill's House Hotel,
115 Meeting Street
Charleston, South Carolina, 29401,
USA**





MEETING AGENDA

| MONDAY MARCH 19, 2007 | | | |
|---|--------------------------------|---|---|
| Registration & Continental Breakfast | | 8:00 am to 9:00 am | |
| Welcome | 9:00 am to 9:15 am | Eric Solon, SWBA President | |
| KEYNOTE SPEAKER 9:15 am to 10:15 am Dr. David Wilson Professor of Biomedical Engineering and Radiology Case Western Reserve University Founder BioInVision, Inc. | | | |
| Session I | Time | Session Chair | Drug Discovery and Development |
| Introduction to Session | 10:15 am to 10:20 am | William Waddel, University of Louisville, US | |
| Speaker IA | 10:20 am to 10:50 am | Alfred Lordi, Quest Pharmaceutical Services, US | Whole-Body Tissue Distribution and Metabolic Profiling of ¹⁴ C-AZT in Fetal and Maternal Tissues |
| Morning Break & Posters | | 10:50 am to 11:20 am | |
| Speaker IB | 11:20 am to 11:50 am | Eric Solon, Quest Pharmaceutical Services, US | Distribution of [¹⁴ C] Dalbavancin in Rabbit Bone and Related Tissues |
| Session Summary | 11:50 am to 12:00 noon | William Waddel, University of Louisville, US | |
| Group Photo | | 12:00-12:15pm | |
| Meeting Luncheon | | 12:15-1:25pm | |
| Session II | Time | Session Chair | Drug Discovery & New Methods |
| Introduction to Session | 1:25 pm to 1:30 pm | Brian Whitby, Covance Labs, UK | |
| Speaker IIA | 1:30 pm to 2:00 pm | Alain Schweitzer, Novartis, Switzerland | Macro-Autoradioluminography of Hard Tissues |
| Speaker IIB | 2:00 pm to 2:30 pm | Marissa Vavrek, Merck, US | Metabolite ID from QWBA Sections Using Advion Nanomate ESI MS |
| Afternoon Break & Posters | | 2:30 pm to 3:00 pm | |
| Speaker IID | 3:00 pm to 3:30 pm | David Wilson, Case Western Reserve University, US | Whole Mouse Cryo-Imaging |
| Speaker IID | 3:30 pm to 4:00 pm | Michael Potchoiba, Pfizer, Groton, Ct, USA | A Quantification Method For Determining the Biodistribution of Tritium Labeled Compounds Using WBAL |
| Session Summary | 4:00 pm to 4:05 pm | Brian Whitby, Covance Labs, UK | |
| Pre-Dinner Cocktails (Bus to Magnolia Plantation @ 5pm) | | 6:00 pm to 7:00 pm | |
| Conference Dinner | (Honors to be Bestowed) | | 7:00 pm to 9:00 pm |



MEETING AGENDA

| TUESDAY MARCH 20, 2007 | | | |
|---|----------------------|--|---|
| Continental Breakfast | | 8:00 am to 9:00 am | |
| Session III | | | |
| | Time | Session Chair | Instrumentation & New Applications |
| Introduction to Session | 9:00 am to 9:05 am | Alfred Lordi, Quest Pharmaceutical Services, US | |
| Speaker IIIA | 9:05 am to 9:35 am | Alain Schweitzer, Novartis | Label-free Molecular Imaging of Whole-body Tissues Sections by Mass Spectrometry |
| Speaker IIIB | 9:35 am to 10:05 am | Ken Koeplinger, Merck, US | QWBA vs 19F Magnetic Resonance Spectroscopy (MRS) for a Fluorinated Nucleoside Drug: Translation of Preclinical Tissue Distribution Studies to the Clinic |
| Morning Break & Posters | | 10:05 am to 10:30 am | |
| Speaker IIIC | 10:30 am to 11:15 am | Tadafumi Kawamoto, Tsurumi University, Japan | Glass Slide Transfer of Whole-Body Section for Histology Examination |
| Speaker IIID | 11:15 am-11:45 am | Helen Minter, Unilever, UK | Skin Penetration and Micro-Autoradiography |
| Session Summary | 11:45 am to 11:50 am | Alfred Lordi, Quest Pharmaceutical Services, US | |
| Meeting Luncheon | | 12:00 noon to 1:00 pm | |
| Session IV | | | |
| | Time | Session Chair | Drug Discovery & New Methods |
| Introduction to Session | 1:00 pm to 1:10 pm | Alain Schweitzer, Novartis, Switzerland | |
| Speaker IVA | 1:10 pm to 1:45 pm | Cynthia Pastukova, Genentech, CA, USA | The effect of immune-complex formation on the biodistribution of [125I] labeled antibody in a mouse model: The use of QWBA in drug discovery |
| Speaker IVB | 1:45 pm to 2:15 pm | Andrew Davis, Cameca, Trumbull, CT, USA | NanoSIMS – Subcellular Isotope Tracer Imaging |
| Poster & Exhibit Viewing & Afternoon Break | | 2:15 pm to 3:00 pm | |
| ISSUES IN PHARMACEUTICAL RESEARCH - OPEN FORUM | 3:00 pm to 5:00 pm | Chairman: Alain Schweitzer, Novartis, Switzerland Panel of Session Chairs | |
| (Dinner is up to individual) | | | |



MEETING AGENDA

| WEDNESDAY, MARCH 21, 2005 | | | |
|---|----------------------|--|--|
| Continental Breakfast | | 8:00 am to 9:00 am | |
| Session V | Time | Session Chair | Regulatory & New Applications continued |
| Introduction to Session | 9:00 am to 9:05 am | Eric Solon, Quest Pharmaceutical Services, US | |
| Speaker VA | 9:05 am to 9:35 am | Brian Whitby, Covance UK | The practicalities of carrying out ADME studies using large molecular weight radiolabelled bio-molecules |
| Speaker VB | 9:35 am to 10:05 am | Birte Hofmann, Bayer HealthCare AG, Germany | Answering Specific Questions Preclinical Questions using WBA |
| Morning Break | | 10:05 am to 10:30 am | |
| Speaker VC | 10:30 am to 11:00 am | Lee Crossman, Schering-Plough, Kenilworth, NJ, USA | Distribution of ¹⁴ C-Florfenicol-Derived Radioactivity to Therapeutically-Targeted Tissues in a Calf Administered a Single Subcutaneous ¹⁴ C-Florfenicol Dose as Resflor NMP (SCH 529752). |
| Speaker VD | 11:00 am to 11:30 am | Alain Schweitzer, Novartis, Basle, Switzerland | Relevance of the Section Thickness Uniformity and Section-to-Section Thickness Reproducibility for Quantification |
| Summary of Session | 11:30 am to 11:45 am | Eric Solon, Quest Pharmaceutical Services, US | |
| Meeting Close Eric Solon | | 11:45 am to 12:00 noon | |

Preparation of Multi-Purpose Fresh-Frozen Sections By Using a New Adhesive Film

Tadafumi Kawamoto, Radioisotope Research Institute, Tsurumi University, School of Dental Medicine, 2-1-3 Tsurumi, Tsurumi-Ku, Yokohama, Japan 230-8501; e-mail address: kawamoto-t@tsurumi-u.ac.jp

I reported on a new approach to preparing thin whole-body sections for histochemistry, immunohistochemistry, and light microscopical autoradiography at this conference in 2001. Cutting procedures were nearly identical to those used in the Ullberg method. However, to better prepare high-quality sections, a new adhesive film and disposable blade were used. This method resulted in 2 μm thick frozen sections that could be examined at high magnification with a light microscope. However, the method is not suitable for routine work because the adhesive film must be prepared before each use. In 2003, I created following adhesive film sheets for routine work. Each film sheet is supported by released paper. Each sheet is cut to match the size of a sample block surface.

1. Cryofilm Type-1 has the strongest adhesive power. It supports a frozen section at approximately -29°C. The section can be immersed in acetone. Sections supported with Cryofilm Type-1 are used for histological staining without toluidine blue, histochemical staining, immunohistochemical staining, and in situ hybridization. The stained section is preserved between the adhesive film and the glass slide.
2. Cryofilm Type-2 has less adhesive power than Cryofilm Type-1. It does not permit dipping in either acetone or xylene. However, it allows histological staining with toluidine blue as well as histochemical staining, immunohistochemical staining, and in situ hybridization.
3. Transfer film is also used for histological staining, histochemical staining, immunohistochemical staining, and in situ hybridization. However, direct transfer of the stained section from the adhesive film to a saline-coated glass slide is possible.
4. LMD film is used to support sections cut by laser microdissection techniques. Collected specimens are used for gene analysis.

I have tried to develop adhesive films usable with a wide range of samples. I have devoted considerable attention to embedding mediums and mounting mediums. Design improvements in disposable blade holders and sample holders have been made. Cryofilm Type-2C is my most recent effort. It is used to support frozen sections at -37°C. It can be immersed in organic solutions such as alcohol, acetone, and xylene. Sections can be placed in a hot chamber (125°C) for more than one hour. Cryofilm Type-2C permits many types of staining including toluidine blue. The film adheres to a frozen block made of a new embedding medium strongly. This is an improvement over conventional embedding mediums such as CMC gel or OCT. Section preparation, staining, and mounting procedures are nearly identical to those described at this conference in 2001. The frozen block is cut with a disposable tungsten carbide blade (Leica TC65). Exposed tissue surface is covered with the adhesive film and then cut into 2~40 μm thick sections. The sections are removed from the cryochamber and instantaneously thawed. They are then immersed in 100% ethanol, fixed with a 4% paraformaldehyde solution (pH 7.4), and stained. The stained sections are sandwiched between the supporting plastic film and the glass slide for permanent preservation. Of course, the new embedding medium and mounting medium are used.

Thin sections from a rabbit thighbone, human bones, and other large samples (adult mice and rats) have been made. Complete thin sections (2 μm) have been prepared from adult rat thighbones, soft tissue, and a whole baby rat. Frozen sections from many other samples are possible. These include plants, fish, insect, grains, and assorted foodstuffs. Sections are applicable to a wide range of research techniques including histological staining, enzyme histochemistry, immunohistochemistry, in situ hybridization, laser microdissection, electron probe microanalysis, and autoradiography of water-soluble materials. Indeed, it is a very versatile and useful tool for a wide spectrum of biological research. At this conference, I will use video and slide presentations to describe my methods and their applications.



MEETING ATTENDEES

| <u>Last Name</u> | <u>First Name</u> | <u>Company name</u> |
|------------------|-------------------|--|
| Anciaux | Kateljine | Janssen Pharmaceutica |
| Brown | Richard | Lablogic Systems, Ltd. |
| Crossman | Lee | Schering-Plough Research Institute |
| Dorenkamp | Claudia | Leica Microsystems Nussloch GmbH |
| Erickson | Jamie | Abbott Bioresearch Center |
| Flood | Dennnis | Novartis Pharmaceuticals |
| Fulton | Jeffrey | Hoffmann-LaRoche, Inc. |
| Geerts | Rita | Janssen Pharmaceutica |
| Hofmann | Birte | Schering AG |
| Kawamoto | Tadafumi | Tsurumi University |
| Knapp | Brian | Covance USA |
| Koeplinger | Ken | Merck Research Labs |
| Korsen | Ann | Leica Microsystems, Inc. |
| LaRocca | Shawn | Leica Microsystems, Inc. |
| Linehan | Stefan | WIL Research Labs, LLC |
| Lordi | Alfred | Quest Pharmaceutical Services |
| Marlowe* | Carolyn | William J. Waddell, Inc. |
| McKown | Jon | biospace USA |
| McNally* | William | SWBA Distinguished Fellow |
| Mehta | Drew | Vibratome Company |
| Minter | Helen | Unilever |
| Partridge | Elizabeth Ann | AstraZeneca |
| Pastukovas | Cinthia | Genetech Inc. |
| Patterson | Andrew | Charles River Laboratories, UK |
| Potchoiba | Michael | Pfizer |
| Press | Randy | Covance USA |
| Rebmann | Nina | QPS |
| Ren | Xiao | sanofi-aventis |
| Schweitzer* | Alain | Novartis Pharma |
| Sved | Dan | WIL Research Labs, LLC |
| Solon | Eric | Quest Pharmaceutical Services |
| Trawick | Dorothy | Array BioPharma |
| Vavrek | Marissa | Merck Research Labs |
| Viot | Delphine | UCB (Belgium) |
| Voller | Thomas | Washington University School of Medicine |
| Wadell* | Bill | University of Louisville |
| Whitby | Brian | Covance Labs Ltd. |
| Wilson | David | Case Western Reserve University |
| Zhen | Ji | sanofi-aventis |
| Zimmer | Manfred | sanofi-aventis |

- Distinguished Fellow of the Society for Whole-Body Autoradiography