
From: Zedler, Barbara
Sent: Sunday, December 12, 2004 6:16 AM
Subject: FW: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne
DSS:

Barb

Barbara K. Zedler, M.D.
Medical Director
Clinical Evaluation
Philip Morris USA
Richmond, VA
barbara.k.zedler@pmusa.com
804.274.2799
cell 804.366.5600
fax 804.274.2391

Confidentiality Notice: This e-mail transmission may contain confidential or legally privileged information that is intended only for the individual or entity named in the e-mail address. If you are not the intended recipient, you are hereby notified that any disclosure, copying, distribution, or reliance upon the contents of this e-mail is strictly prohibited.

If you have received this e-mail transmission in error, please reply to the sender, so that we can arrange for proper delivery, and then please delete the message from your inbox. Thank you.

-----Original Message-----

From: Zedler, Barbara
Sent: Thursday, May 27, 2004 4:23 PM
To: Roethig, Hans
Cc: Zedler, Barbara
Subject: FW: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

It would be most helpful for PMUSA to have a single, comprehensive topography working group with knowledge, oversight, and review of all topography initiatives/studies being conducted within PMUSA in order to communicate clearly and uniformly about this topic.

Barb

Barbara K. Zedler, M.D.
Director, Clinical Operations
Clinical Evaluation
Worldwide Scientific Affairs
Philip Morris USA
Richmond, VA

1/7/2005

PM3008423423

3008423423

3008423423

barbara.k.zedler@pmusa.com
804.274.2799
cell 804.366.5600

-----Original Message-----

From: Sarkar, Mohamadi
Sent: Thursday, May 27, 2004 4:16 PM
To: Werley, Michael S; Mendes, Paul; Tricker, Anthony; Zedler, Barbara; Reininghaus, Wolf; Dempsey, Ruth
Subject: RE: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

No, no, no!!! Our study does not use the average topography parameters (which is the reason for this being such a unique experiment). We are using varying parameters over each puff, hence the delay since the smoke machine cannot handle dynamic smoking regimen.

Mohamadi

-----Original Message-----

From: Werley, Michael S
Sent: Thursday, May 27, 2004 4:04 PM
To: Mendes, Paul; Sarkar, Mohamadi; Tricker, Anthony; Zedler, Barbara; Reininghaus, Wolf; Dempsey, Ruth
Subject: RE: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

Dear Everybody,

We have done such a study with the help of PMRL, that is, programming a rotary smoking machines with topography parameters determined

from the human clinical studies (K Short-term). Your project sounds similar to what we have already done. This data should be available

at the meeting in a couple of weeks. We needed to do this to establish nicotine deliveries under topography regimes, using the smoking

machine as a kind of simulated human. Once we know how to hit the nicotine target value (from the clinical study), we can then collect

smoke condensate to test in the REST studies, comparing condensates from K-Series, MUL, and ML. I hope that adds some clarification.

-----Original Message-----

From: Mendes, Paul
Sent: Thursday, May 27, 2004 3:18 PM
To: Sarkar, Mohamadi; Tricker, Anthony; Zedler, Barbara; Werley, Michael S; Reininghaus, Wolf; Dempsey, Ruth
Subject: RE: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

Mohamadi, thanks for the clarification. Sorry Tony, I wasn't aware of this.

Best regards,

Paul

-----Original Message-----

From: Sarkar, Mohamadi
Sent: Thursday, May 27, 2004 2:43 PM
To: Mendes, Paul
Cc: Tricker, Anthony

1/7/2005

PM3008423424

3008423424

3008423424

Subject: RE: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

Dear Tony,

I found out indirectly about your request. As you very well know that I have been analyzing the topography data from our clinical studies and trying to make sense of the relationship with exposure. The key experiment that Hans was referring to is not what is described by Paul but is another research project that Hans and I have initiated in collaboration with David Self. Basically this involves measuring nicotine yields under the variable smoking regimen as observed under human conditions. This experiment has not even been started due to their inability to program the smoke machine. The availability of results is still undetermined which was the reason Hans was suggesting that this workshop be postponed till we get those results back as well as know more from the Plowshare device validation studies. I can send you the experimental details and project sheet if you are interested.

Best Regards,

Mohamadi

-----Original Message-----

From: Mendes, Paul

Sent: Thursday, May 27, 2004 2:03 PM

To: Sarkar, Mohamadi

Subject: FW: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

-----Original Message-----

From: Mendes, Paul

Sent: Thursday, May 27, 2004 1:28 PM

To: Tricker, Anthony; Zedler, Barbara

Cc: Werley, Michael S; Reininghaus, Wolf; Dempsey, Ruth

Subject: RE: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

Dear Tony,

The Key Experiment that Hans is referring to is definitely not a clinical study. I believe it is the experiment that Mike is currently running at PMRL, setting puff volumes, puff durations, interpuff intervals to achieve the nicotine values found in clinical studies, that he meant. To the best of my knowledge the results will be available next week or the week thereafter. Hope this is of help.

Best Regards,

Paul

-----Original Message-----

From: Tricker, Anthony

Sent: Thursday, May 27, 2004 11:36 AM

To: Mendes, Paul; Zedler, Barbara

Cc: Werley, Michael S; Reininghaus, Wolf; Dempsey, Ruth; Tricker, Anthony

Subject: FW: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

Dear Barbara, Dear Paul,

I have just heard that Hans is away and cannot address the following email that I sent him a few minutes ago. As you appreciate it is important that we move forward in trying to understand the limitations in using Plowshare, hence it is very important that something is said about the "key experiment" that Hans refers to at the Cologne meeting. However, at present, no one here has a clue what this experiment is. Can you make sure that Mike has some information on Hans's "key experiment" so that this can be presented at the meeting.

1/7/2005

PM3008423425

3008423425

3008423425

With best regards,
Tony

-----Original Message-----

From: Tricker, Anthony
Sent: Thursday, 27. May 2004 17:21
To: Roethig, Hans
Cc: Dempsey, Ruth; Reininghaus, Wolf; Lewis, Jane Y.; Tricker, Anthony; Werley, Michael S
Subject: FW: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

Dear Hans,

It is unfortunate that you cannot send anyone from your group to present a summary on the US clinical evaluations experience with Plowshare; however, I have no doubt that Mike Werley can do a marvelous job on presenting the supplied material. One thing which remains unclear - will this also describe the "key experiment" you refer to as know one here knows what you are referring to. If we are to plan for future studies and evaluations to justify the continued use of Plowshare, we also need to know what is currently in progress or planned to avoid unnecessary duplication of work.

With best regards,
Tony

-----Original Message-----

From: Roethig, Hans
Sent: Thursday, 20. May 2004 21:17
To: Dempsey, Ruth; Tricker, Anthony
Cc: Reininghaus, Wolf; Werley, Michael S; Lewis, Jane Y.; Sarkar, Mohamadi; Zedler, Barbara; Mendes, Paul
Subject: RE: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

Dear Ruth, dear Wolf, dear Tony,
Mike Werley has kindly agreed to present the information we have. Mohamadi Sarkar will give him the information next week. In addition we have a final draft of the ACCORD JLI Short Term Exposure Study report, which includes information on topography. Barbara Zedler will send the topography information and interpretation to Mike next week. I hope this will be satisfactory to the group.

My proposal to postpone the workshop was based on the intention to base decisions on important facts to come. In the discussions about topography publications in the former WSA organization Rick and his staff had requested to postpone any publications until we had these results.

Best regards
Hans

-----Original Message-----

From: Dempsey, Ruth
Sent: Thursday, May 20, 2004 2:48 AM
To: Roethig, Hans; Tricker, Anthony
Cc: Reininghaus, Wolf
Subject: RE: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

Hans,

I really do not think that we should postpone this workshop. Whilst I understand if you really cannot spare someone for those dates we really would like to have some of your data to include in the discussions. I am very curious, for example, on the expected contribution from the "key experiment" that you describe. Just understanding how you arrived at concluding that this latest experiment was required would be an extremely valuable contribution to the workshop. I believe that other individuals who could not attend have either sent representatives knowledgeable in their area or are sending presentation material independently. I wonder if you could manage the latter? Tony, at least, should be familiar with all the background and should be able to

1/7/2005

PM3008423426

3008423426

3008423426

represent such a presentation fairly faithfully since I presume that you have included Tony regularly in discussions on such research areas in the clinical evaluations group.

I am sure that with a little cooperation we can work something out which will be satisfactory.

with many thanks and kind regards,
Ruth

-----Original Message-----

From: Roethig, Hans
Sent: Wednesday, 19. May 2004 17:55
To: Tricker, Anthony
Cc: Reininghaus, Wolf; Dempsey, Ruth
Subject: RE: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

If the dates could be changed, we could send someone.
We have done a lot with topography data, but we are lacking results from a key experiment.
I would postpone the workshop until autumn this year.
Best regards
Hans

-----Original Message-----

From: Tricker, Anthony
Sent: Wednesday, May 19, 2004 10:55 AM
To: Roethig, Hans
Cc: Reininghaus, Wolf; Dempsey, Ruth
Subject: RE: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne
Importance: High

Dear Hans,
You are absolutely right when you state that a clinical exposure expert is lacking at this meeting. Do you have any recommendation of someone who could attend this meeting from the clinical evaluations group in Richmond? It would be very beneficial to all parties if this expertise was present and could present a summary of the key findings and learning's from the studies you have performed.

Looking forward to hearing from you in the near future.
With best regards,
Tony

-----Original Message-----

From: Roethig, Hans
Sent: Friday, 14. May 2004 21:52
To: Reininghaus, Wolf; Lindner, Dirk; Martin Leroy, Claire; Rigaux, Michel; Sanders, Edward; Schneider, Jean-Claude; Schorp, Matthias; Tricker, Anthony; Counts, Mary Ellen; Lipowicz, Peter J.; Livermore, Andrew; Koller, Debbie; Werley, Michael S; Dwyer, R.W. (Bill); Gogova, Maria

Cc: Solana, Rick P.; Lewis, Jane Y.; Kobal, Gerd; Dempsey, Ruth; Walk, Roger A.; Alonso, Hector; Patskan, George J.
Subject: RE: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

Dear Wolf,
Maria Gogova is no longer in Clinical Evaluation.
I think without a clinical topography expert and a clinical exposure expert you will lack substantial knowledge for this workshop.

Colleagues in Ashland have done a great job and evaluated the topography device, which we have used in our clinical studies, on smoking machines.

1/7/2005

PM3008423427

3008423427

3008423427

The outcome was that the device has several problems, so the data have to be regarded very carefully and have serious limitations, in particular for absolute values.

Only a substantially improved topography device may allow us in the future to generate more accurate data, which may help us to set up smoking machines comparable to how smokers puff.

Best regards
Hans

-----Original Message-----

From: Reininghaus, Wolf

Sent: Friday, May 14, 2004 1:32 PM

To: Lindner, Dirk; Martin Leroy, Claire; Rigaux, Michel; Sanders, Edward; Schneider, Jean-Claude; Schorp, Matthias; Tricker, Anthony; Counts, Mary Ellen; Lipowicz, Peter J.; Livermore, Andrew; Koller, Debbie; Werley, Michael S; Dwyer, R.W. (Bill); Gogova, Maria

Cc: Solana, Rick P.; Lewis, Jane Y.; Kobal, Gerd; Dempsey, Ruth; Roethig, Hans; Walk, Roger A.; Alonso, Hector; Patskan, George J.

Subject: REST - DESIGN Workshop, June 7th and 8th, 2004, Cologne

Dear colleagues,

I would like to come back to Tony's suggestion and organize a workshop on smoking topography and machine smoking parameters as they relate to the design of REST studies.

Since several groups at PMI and PMUSA are working in parallel and under different aspects on related topics, an exchange of data, results and opinions might bring us forward.

The purpose of this E-Mail is to fix a meeting date, location, the participation, to agree on the meeting's basic objectives and to solicitate commitments for contributions.

As a starter, I propose the following structure for the workshop:

REST – DESIGN Workshop

General Objective:

Develop a concept / strategy for (PMI) REST studies. (What topography and machine parameters determine smoke composition and exposure and should be considered in a REST study design?)

Topics for short presentations / summaries / overviews:

1. Review of available data on machine cigarette yields, topography data, and clinical exposure doses (Biomarkers): Which reliable data are available, where?

2. Correlations between smoke constituents yields obtained under different machine smoking conditions:

How can we use relations (e. g. Burn time, Multiple Regression (Case), Coal Volume etc) to evaluate, refine, understand and predict yields?

3. Assessment and interpretation of human smoking topography:
What do we know about human smoking parameters (their confidence limits, identification of and dealing with error, pattern in human smoking habits) and which are important for the estimation of exposure?

1/7/2005

PM3008423428

3008423428

3008423428

4. Assessment and interpretation of clinical exposure biomarker data:
How can we use clinical biomarker data (e. g. dealing with background variability and outliers, averaging data, correlation of uptake and yield) to define the smoking parameters in REST studies?

Topics for discussions and results:

- Proposed REST study design (PMI)
- Comparison of PMI / PMUSA approaches

Logistics:

Date : June 7th, 2004 12:00 – 18:00

June 8 th, 2004 8:00 – 18:00

in Cologne, Germany

List of possible participants:

PMI: DirkLinder
Claire Martin

Michel Rigaux

Ted Sanders

Jean-Claude Schneider

Mathias Schorp

Anthony Tricker

PMUSA: Mary Ellen Counts

Bill Dwyer

Peter Lipowicz

Andrew Livermore

D. Koller

Mike Werley

Maria Gogova

PMRL-G: Birgit Gerstenberg

Willy Gomm

Falk Radtke

Wolf Reininghaus

Klaus Rustemeier

1/7/2005

PM3008423429

3008423429

3008423429

H. Schaffernicht

H.-Joerg Urban

The remaining time to organize this meeting is short. I therefore ask for your fast response to the following questions:

1. will you be able to participate?
2. do you propose additional topics?
3. what would you present? (propose short title and duration)
4. do you have other suggestions? (topics, participants,...)

I would appreciate your fast response very much. Best regards

Wolf Reininghaus

eMail wolf.reininghaus@pmintl.com

□□,@B

1/7/2005

PM3008423430

3008423430

3008423430