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**Id :** i.m.07a6d3e68dbf9bfa5266068e31a47016  
**CN :** SQ1ED00099305  
**Date :** Friday, March 10, 2000 3:19:00 PM GMT  
**From :** Brecher Martin M  
**To :** Holdsworth Debbie D; Jones Martin AM - PHMS; Westhead Emma EK  
**Cc :** Aked Dominic DM  
**Subject :** RE: weight gain  
**Custodians :** Jones, Martin

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From: Brecher Martin M  
Sent: 3/10/2000 6:49:00 PM  
To: Holdsworth Debbie D; Jones Martin AM - PHMS; Westhead Emma EK  
CC: Aked Dominic DM  
BCC:  
Subject: RE: weight gain

Emma,

Were all the patients represented on the graph on seroquel alone or some receiving other neuroleptics or mood stabilizers. If patients were on these concomitant meds which cause weight gain, we should re-analyze with those patients excluded.

The data depicted is more accurately characterized by a minimal weight gain statement, but weight neutral is defensible. Assuming the graph depicts patients on seroquel alone we should compare our data with that supporting ziprasidone's weight neutral claim.

Since weight gain will be an important point of differentiation for years to come we should have broad consensus among biostats, clinical, marketing and regulatory.

Martin

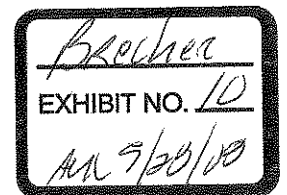
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>From: Westhead Emma EK  
>Sent: Wednesday, March 08, 2000 5:58 AM  
>To: Brecher Martin M; Holdsworth Debbie D; Jones Martin AM - PHMS  
>Cc: Aked Dominic DM  
>Subject: RE: weight gain  
>Importance: High

>

>Please find attached a revised IIIb OLE weight graph (table format and plot).

>



>If we consider up to a year not all of the lower bounds of the CIs are below zero. 2 of the lower bounds are at 0.87 and 0.55.

>After one year, only the final data point (which is just 9 patients) includes zero in the CI.

>

>I think this data is again useful to defend minimal weight gain, but don't think it can support the weight neutral hypothesis suggested below.

>

>Please let me know any comments or any amendments to be made.

>

>Regards,

>Emma

>

><<File: Td9\_t1.doc>><<File: IIIb OLE weight graph with CI.doc>>

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> From: Jones Martin AM - PHMS

> Sent: 07 March 2000 17:39

> To: Brecher Martin M; Holdsworth Debbie D

> Cc: Emma Westhead; Aked Dominic DM

> Subject: RE: weight gain

>

> Debbie/Martin

>

> Yesterday, I asked Emma to progress this analysis ASAP.

>

> Martin

>

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> From: Holdsworth Debbie D

> Sent: 07 March 2000 10:22

> To: Jones Martin AM - PHMS; Brecher Martin M

> Subject: FW: weight gain

>

> fyi

>

> From: Aked Dominic DM

> Sent: 04 March 2000 00:22

> To: Westhead Emma EK

> Cc: Litherland Steve S; Holdsworth Debbie D

> Subject: weight gain

- >
- > Hi Emma
- >
- > I'm exploring my understanding of what we can support on weight gain. Whilst the mean weight changes summaries the data it says nothing about the variation.
- >
- > If .....the upper and lower limits of 95% CI for weight are sufficiently wide (with the lower limit consistently below zero) to suggest that the mean weight gain of 2-3 kg is unlikely to be significantly different from 'zero'.....
- >
- > See attached figure.....
- >
- > Could we defend 'weight neutral' ....meaning not either weight gain or weight loss?
- >
- > I'm wondering whether we could define 'weight neutral' as the lower limit of the 95% CI being consistently below zero. This would clarify what we meant by the term and thereby provide both a scientific and promotional defence. In doing this we would also 'spoil' one of Pfizer's likely key promotional claims!
- >
- > Any thoughts
- >
- > Kind regards
- >
- > Dom
- >
- > <<File: hypothetical weight gain.ppt>>
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