

(cont. from page. start )

Purpose: To prepare TRAVATAN BAC Free, 0.004% with pH 5.5, 6.5 and with 85% of zinc, boric acid for PET  
H. Smith 5-3-05

**Experimental:**

Equipment:

Mettler AX 205 DR Analytical Balance, Asset No. 515840, Rm. B134 (referred to as Balance 1)  
Mettler PG 5002 Balance, Asset No. 101265, Rm. B134 (referred to as Balance 2)  
Orion 525A+ pH Meter, Asset No. A2309, Rm. B134  
The Advanced Micro-Osmometer Model 3300, Asset no. 00103390, Rm. B132  
Stir Plates, Rm. B134  
Millipore 0.22 µm Filter Unit (1000 mL)

Formulations:

FID	Batch	Description
109032	05-40452	TRAVATAN BAC Free, 0.004% - pH 5.5
109033	05-40453	TRAVATAN BAC Free, 0.004% - pH 6.5
109034	05-40454	TRAVATAN BAC Free, 0.004% - pH 5.5 w/ 85% of zinc, boric acid

**Compounding Procedures:** (see attached batch records with adjunctive data)

*Handwritten initials and date: H. Smith 5-4-05*

- A. FID 109032 Batch 05-40452 TRAVATAN BAC Free, 0.004% - pH 5.5
  1. Tare weighed a clean glass beaker with a stir bar. (Balance 2) *AS 5-3-05*
  2. Removed the stir bar. Weighed and added the batch quantity of HCO-40 directly in the beaker. (Balance 2) *AS 5-3-05*
  3. Added about 50 % of batch quantity of purified water. *AS 5-3-05*
  4. Stirred for at least 15 minutes until HCO-40 is dissolved. *AS 5-3-05*
  5. Weighed and added the batch quantity of the following excipients to the beaker: boric acid, sorbitol, propylene glycol, and zinc chloride. Stirred the solution to dissolve these ingredients. (Balance 1) *AS 5-3-05*
  6. Qsed to 95% of batch weight with purified water. (Balance 2) *AS 5-3-05*
  7. Measured the pH. Initial pH = 4.010 *AS 5-3-05*
  8. Adjusted pH if necessary.
    - Amount of 1N HCL added = N/A *AS 5-3-05*
    - Amount of 1N NaOH added = 0.40g
  9. Weighed the batch quantity of AL-06221 using a spatula. (Balance 1) *AS 5-3-05*  
The spatula was placed in the bottle with the end containing AL-06221 immersed in the solution.
  10. Covered the opening with parafilm to minimize evaporation. Stirred overnight to dissolve the AL-06221.  
Stirring Start time and date: 3:15 pm AS 5-3-05  
Stirring Stop time and date: 9:00 am AS 5-4-05
  11. Qsed to 100% of batch weight with purified water. (Balance 2) Final pH = 5.573 *AS 5-4-05*
  12. Measured Osmolality. Osmolality = 283 mOsm/L *AS 5-4-05*

ADJUNCTIVE DATA *AS 6-21-05*

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Verified By : <i>BKARBYA</i>	Date <i>6-24-05</i>	Entered By : <i>H. Smith</i>	Date <i>6-21-05</i>
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B. FID 109033 Batch 05-40453 TRAVATAN BAC Free, 0.004% - pH 6.5

1. Tare weighed a clean glass beaker with a stir bar. (Balance 2) AS 5-3-05
2. Removed the stir bar. Weighed and added the batch quantity of HCO-40 directly in the beaker. (Balance 2) AS 5-3-05
3. Added about 50 % of batch quantity of purified water. AS 5-3-05
4. Stirred for at least 15 minutes until HCO-40 is dissolved. AS 5-3-05
5. Weighed and added the batch quantity of the following excipients to the beaker: boric acid, sorbitol, propylene glycol, and zinc chloride. Stirred the solution to dissolve these ingredients. (Balance 1) AS 5-3-05
6. Qsed to 95% of batch weight with purified water. (Balance 2) AS 5-3-05
7. Measured the pH. Initial pH = 4.076 AS 5-3-05
8. Adjusted pH if necessary.  
Amount of 1N HCL added = N/A AS 5-3-05  
Amount of 1N NaOH added = 1.49g
9. Weighed the batch quantity of AL-06221 using a spatula. (Balance 1) AS 5-3-05  
The spatula was placed in the bottle with the end containing AL-06221 immersed in the solution.  
10. Covered the opening with parafilm to minimize evaporation. Stirred overnight to dissolve the AL-06221.  
Stirring Start time and date: 3:15 pm AS 5-3-05  
Stirring Stop time and date: 9:00 am AS 5-4-05
11. Qsed to 100% of batch weight with purified water. (Balance 2) Final pH = 6.553 (2) AS 5-4-05
12. Measured Osmolality. Osmolality = 278 mOsm/kg AS 5-4-05

C. FID 109034 Batch 05-40454 TRAVATAN BAC Free, 0.004% - pH 5.5 w/ 85% of zinc, boric acid

1. Tare weighed a clean glass beaker with a stir bar. (Balance 2) AS 5-3-05
2. Removed the stir bar. Weighed and added the batch quantity of HCO-40 directly in the beaker. (Balance 2) AS 5-3-05
3. Added about 50 % of batch quantity of purified water. AS 5-3-05
4. Stirred for at least 15 minutes until HCO-40 is dissolved. AS 5-3-05
5. Weighed and added the batch quantity of the following excipients to the beaker: boric acid, sorbitol, propylene glycol, and zinc chloride. Stirred the solution to dissolve these ingredients. (Balance 1) AS 5-3-05
6. Qsed to 95% of batch weight with purified water. (Balance 2) AS 5-3-05
7. Measured the pH. Initial pH = 4.103 AS 5-3-05
8. Adjusted pH if necessary.  
Amount of 1N HCL added = N/A  
Amount of 1N NaOH added = 0.37g AS 5-3-05
9. Weighed the batch quantity of AL-06221 using a spatula. (Balance 1) AS 5-3-05  
The spatula was placed in the bottle with the end containing AL-06221 immersed in the solution.
10. Covered the opening with parafilm to minimize evaporation. Stirred overnight to dissolve the AL-06221.  
Stirring Start time and date: 3:15 pm AS 5-3-05  
Stirring Stop time and date: 9:00 am AS 5-4-05
11. Qsed to 100% of batch weight with purified water. (Balance 2) Final pH = 5.574 AS 5-4-05
12. Measured Osmolality. Osmolality = 257 mOsm/kg AS 5-4-05

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Verified By :

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BOOK NO: 11781

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TITLE TRAVATAN BAC Free, 0.004% Formulation for AET

PROJECT # 225719

(cont. from page 20) AS 5-1-05

Filter Batches:

Filter each batch using a 250ml Millipore Filter Unit 0.22um  
Lot H4KN90620 expires sep 2007

Lot Reports: AS 5-4-05

Print lot reports for each lot.

ADJUNCTIVE DATA AS 6-21-05

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PET Submission: AS 5-4-05

Submit lots 05-40452-1, 05-40453-1, and 05-40454-1 to microbiology for  
PET, USP LIMS submission ID: 100082058

Analytical Request (Chemistry): AS 5-5-05

Submit each lot tested in microbiology to analytical chemistry for the following test:  
sample of AS 5-5-05

Boric acid 1.0% Assay

Zinc 0.0025% Assay

PET Results: AS 6-21-05

All lots submitted for PET, USP pass USP requirements. Sample Results report  
with adjunctive data.

Analytical Chemistry Results: AS 6-21-05

see adjunctive data for all analytical chemistry results.

N/A AS 6-21-05

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Verified By :

Bhanna

Date

6-24-05

Entered By :

[Signature]

Date

6-21-05