

**Assessment of dermal absorption of aluminium from a representative antiperspirant formulation using a [<sup>26</sup>Al]Al microtracer approach – a follow-up study in humans**

Rianne de Ligt<sup>1</sup>, Joost Westerhout<sup>2</sup>, Dimitri Grossouw<sup>1</sup>, Thomas P. Buters<sup>3</sup>, Robert Rissmann<sup>3</sup>, Jacobus Burggraaf<sup>3</sup>, Albert D. Windhorst<sup>4</sup>, Sarah Tozer<sup>5</sup>, Gerlinde Pappa<sup>6</sup>, Brian Wall<sup>7</sup>, Dagmar Bury<sup>8</sup>, David R. Mason<sup>9</sup>, and Wouter H.J. Vaes<sup>1</sup>

Table S1 – Overview of nominal doses applied (parts A and B)

| <b>Part – Treatment</b> | <b>Amount</b> | <b>Concentration</b> | <b>Nominal dose</b> | <b>Nominal dose of [<sup>26</sup>Al]</b> |
|-------------------------|---------------|----------------------|---------------------|--|
| A – Topical (~2500 Bq)  | 1.5 g         | 1797 Bq/g            | 2695 Bq             | 3730317 pg                               |
| A – IV (cohort 1)       | 5 mL          | 0.017 Bq/mL          | 0.086 Bq            | 120 pg                                   |
| A – IV (cohort 2)       | 5 mL          | 0.014 Bq/mL          | 0.072 Bq            | 100 pg                                   |
| B – Topical (~1 Bq)     | 1.5 g         | 0.76 Bq/g            | 1.14 Bq             | 1573 pg                                  |

Subjects 01-06 were included in Part A, where cohort 1 was comprised of subjects 01, 03, 04 and 05 and cohort 2 comprised of subjects 02 and 06, subjects 07-12 were included in Part B

**Assessment of dermal absorption of aluminium from a representative antiperspirant formulation using a [<sup>26</sup>Al]Al microtracer approach – a follow-up study in humans**

Rianne de Ligt<sup>1</sup>, Joost Westerhout<sup>2</sup>, Dimitri Grossouw<sup>1</sup>, Thomas P. Buters<sup>3</sup>, Robert Rissmann<sup>3</sup>, Jacobus Burggraaf<sup>3</sup>, Albert D. Windhorst<sup>4</sup>, Sarah Tozer<sup>5</sup>, Gerlinde Pappa<sup>6</sup>, Brian Wall<sup>7</sup>, Dagmar Bury<sup>8</sup>, David R. Mason<sup>9</sup>, and Wouter H.J. Vaes<sup>1</sup>

Table S2 – Summary of emergent adverse events by relationship, intensity and system organ class

| <b>Relationship</b> | <b>Intensity</b> | <b>System Organ Class (number)</b>  |
|---------------------|------------------|---|
| Probably            | Moderate         | Infusion site pain (2)  |
|                     | Mild             | Dermatitis contact (3); Skin abrasion (2); Application site erythema (1); Dizziness (1); Nausea (1); Pruritus (1) |
| Possibly            | Mild             | Paraesthesia (1)  |
| Unrelated           | Moderate         | Influenza like illness (1); Musculoskeletal pain (1)  |
|                     | Mild             | Headache (14); General disorders (4)  |

**Assessment of dermal absorption of aluminium from a representative antiperspirant formulation using a [<sup>26</sup>Al]Al microtracer approach – a follow-up study in humans**

Rianne de Ligt<sup>1</sup>, Joost Westerhout<sup>2</sup>, Dimitri Grossouw<sup>1</sup>, Thomas P. Buters<sup>3</sup>, Robert Rissmann<sup>3</sup>, Jacobus Burggraaf<sup>3</sup>, Albert D. Windhorst<sup>4</sup>, Sarah Tozer<sup>5</sup>, Gerlinde Pappa<sup>6</sup>, Brian Wall<sup>7</sup>, Dagmar Bury<sup>8</sup>, David R. Mason<sup>9</sup>, and Wouter H.J. Vaes<sup>1</sup>

Table S3 – Part B. Averaged amount of [<sup>26</sup>Al] recovered from the tape strips and skin biopsies, normalized for the amount recovered after 20 minutes of application.

| Tape strip number(s)               | Recovery in % of dose (normalized on t=20 min) |      |      |       |
|------------------------------------|--|------|------|-------|
|                                    | t=20 min                                       | t=1h | t=6h | t=24h |
| 1                                  | 70%  | 44%  | 7.0% | 0.6%  |
| 2                                  | 13%  | 11%  | 1.0% | 0.2%  |
| 3                                  | 4.5%   | 3.8% | 0.9% | 0.2%  |
| 4                                  | 3.4%   | 2.0% | 0.3% | 0.1%  |
| 5                                  | 2.0%   | 1.4% | 0.3% | 0.1%  |
| 6                                  | 2.3%   | 0.8% | 0.2% | 0.04% |
| 7                                  | 1.0%   | 1.4% | 0.8% | 0.1%  |
| 8                                  | 0.9%   | 1.0% | 0.7% | 0.1%  |
| 9                                  | 0.9%   | 0.9% | 0.4% | 0.1%  |
| 10                                 | 0.9%   | 0.9% | 0.4% | 0.1%  |
| 11-20                              | 0.6%   | 0.5% | 0.1% | 0.03% |
| 21-30                              | 0.5%   | 0.3% | 0.1% | 0.02% |
| 31-40                              | 0.3%   | 0.1% | 0.1% | 0.02% |
| Total                              | 100%   | 68%  | 12%  | 1.6%  |
| Skin biopsy (n=4, 2 samples <LLOQ) | n.a.   | n.a. | n.a. | 0.08% |

n.a. not applicable