

### Disposition of Radioactivity 24 Hours Following 5 Days of Oral Gavage Administration of 1 mg/kg [ $^{14}\text{C}$ ]L-BMAA to Male Harlan Sprague Dawley Rats (Study F)<sup>a</sup>

#### Dose Recovered in Excreta (%)

| End of Collection Period (h) | Urine CPDE <sup>b</sup> | Cage Rinse CPDE   | Feces CPDE       | Volatile Organics CPDE | CO <sub>2</sub> CPDE | GI Content        | Total CPDE |
|------------------------------|-------------------------|-------------------|------------------|------------------------|----------------------|-------------------|------------|
| 4                            | 0.226 $\pm$ 0.161       | 0.137 $\pm$ 0.082 | —                | 0.017 $\pm$ 0.082      | 27.1 $\pm$ 5.95      | —                 | 27.5       |
| 8                            | 0.712 $\pm$ 0.133       | 0.227 $\pm$ 0.095 | —                | 0.034 $\pm$ 0.095      | 41.6 $\pm$ 6.38      | —                 | 42.6       |
| 12                           | 0.899 $\pm$ 0.112       | 0.305 $\pm$ 0.118 | —                | 0.054 $\pm$ 0.118      | 47.9 $\pm$ 5.97      | —                 | 49.2       |
| 24                           | 1.23 $\pm$ 0.066        | 0.340 $\pm$ 0.117 | 1.16 $\pm$ 0.296 | 0.073 $\pm$ 0.117      | 52.8 $\pm$ 4.98      | —                 | 55.6       |
| 48                           | 2.40 $\pm$ 0.271        | 0.426 $\pm$ 0.134 | 1.78 $\pm$ 0.368 | 0.048 $\pm$ 0.134      | 54.7 $\pm$ 3.08      | —                 | 59.3       |
| 72                           | 3.54 $\pm$ 0.175        | 0.547 $\pm$ 0.087 | 3.02 $\pm$ 0.821 | 0.038 $\pm$ 0.087      | 53.3 $\pm$ 3.75      | —                 | 60.4       |
| 96                           | 4.74 $\pm$ 0.301        | 0.709 $\pm$ 0.128 | 4.46 $\pm$ 1.41  | 0.034 $\pm$ 0.128      | 53.2 $\pm$ 3.83      | —                 | 63.2       |
| 120                          | 6.07 $\pm$ 0.327        | 1.072 $\pm$ 0.078 | 6.22 $\pm$ 1.34  | 0.032 $\pm$ 0.078      | 52.6 $\pm$ 3.48      | 0.455 $\pm$ 0.362 | 66.5       |

#### Disposition in Tissues

| Tissue               | nmol-eq/g Mean | nmol-eq/g SD | TBR Mean | TBR SD | % Recovery Mean | % Recovery SD |
|----------------------|----------------|--------------|----------|--------|-----------------|---------------|
| Blood <sup>d</sup>   | 3.30           | 0.293        | unity    | —      | 0.568           | 0.036         |
| Adipose <sup>d</sup> | 2.86           | 0.671        | 0.869    | 0.207  | 0.466           | 0.105         |
| Muscle <sup>d</sup>  | 2.73           | 0.466        | 0.825    | 0.110  | 2.56            | 0.379         |
| Skin <sup>d</sup>    | 5.25           | 0.792        | 1.59     | 0.196  | 2.32            | 0.290         |
| Brain                | 1.77           | 0.261        | 0.539    | 0.046  | 0.027           | 0.004         |
| Heart                | 3.63           | 1.059        | 1.10     | 0.203  | 0.027           | 0.009         |
| Kidneys              | 7.93           | 0.462        | 2.43     | 0.101  | 0.122           | 0.007         |
| Liver                | 15.00          | 1.823        | 4.59     | 0.270  | 1.20            | 0.120         |
| Lung                 | 4.35           | 0.340        | 1.33     | 0.052  | 0.049           | 0.004         |
| Spleen               | 4.34           | 0.527        | 1.34     | 0.203  | 0.025           | 0.003         |
| Adrenals             | 9.96           | 2.095        | 3.03     | 0.659  | 0.003           | 0.001         |
| Thymus               | 5.54           | 0.921        | 1.72     | 0.380  | 0.025           | 0.008         |
| Thyroid              | 10.53          | 6.858        | 4.02     | 1.151  | 0.000           | 0.000         |
| Urinary Bladder      | 5.61           | 1.268        | 1.71     | 0.342  | 0.003           | 0.000         |
| Pancreas             | 4.99           | 1.343        | 1.55     | 0.443  | 0.031           | 0.010         |
| Testes               | 2.10           | 0.316        | 0.640    | 0.050  | 0.059           | 0.013         |
| Cecum                | 4.51           | 0.751        | 1.38     | 0.193  | 0.018           | 0.004         |
| Large Intestine      | 4.53           | 0.798        | 1.39     | 0.217  | 0.041           | 0.006         |
| Small Intestine      | 5.36           | 0.616        | 1.64     | 0.138  | 0.159           | 0.033         |
| Stomach              | 4.39           | 0.607        | 1.34     | 0.144  | 0.054           | 0.005         |

#### Disposition Summary (Dose Recovered [%])

| Sample                 | % Recovered Mean | % Recovered SD |
|------------------------|------------------|----------------|
| Urine + Cage Rinse     | 7.14             | 0.328          |
| Urine                  | 6.07             | 0.405          |
| Cage Rinse             | 1.07             | 0.078          |
| Feces                  | 6.22             | 1.35           |
| Volatile Organics      | 0.032            | 0.005          |
| CO <sub>2</sub>        | 52.6             | 3.48           |
| GI Content             | 0.455            | 0.362          |
| Carcass                | 0.277            | 0.284          |
| Tissues                | 8.04             | 0.675          |
| <b>Total Recovered</b> | <b>74.8</b>      | <b>3.52</b>    |

<sup>a</sup>All values expressed as mean  $\pm$  standard deviation (SD) (N = 5). The target dose was 1 mg L-BMAA/kg. The actual dose delivered was 0.994  $\pm$  0.017 mg/kg (41.4  $\pm$  0.75  $\mu\text{Ci}$ ).

<sup>b</sup>CPDE = Cumulative percent dose excreted

<sup>c</sup>120 hours (h) urine collection includes urine present in the urinary bladder at study termination.

<sup>d</sup>Tissue weights for the dispersed tissues were calculated using the following percentages of body weight: adipose 7.0%, blood 7.4%, muscle 40.4%, and skin 19% (International Life Sciences Institute, 1994).

TBR = Tissue to blood ratio. Rats were dosed daily for 5 days with sacrifice 24 hours post last dose.