

Supplemental Figure 1. Four patients (from left to right: patient \#1 in arm A, and patient \#2, \#7, \#16 in arm B) with fulminant hepatic metastases.

Supplemental Table 1. Inclusion and exclusion criteria

## Inclusion Criteria

- Written informed consent.
- Patients of either gender, aged $\geqslant 18$ years.
- Histologically confirmed diagnosis of Metastatic, well-differentiated neuroendocrine tumor.
- A diagnostic computed tomography (CT) or magnetic resonance imaging (MRI) of the tumor region within the previous 6 months prior to dosing day is available.
- At least 1 measurable lesion based on RECIST v1.1.
- Blood test results as follows (White blood cell: $\geqslant 3 * 10^{\wedge} 9 / L$, Hemoglobin: $\geqslant 8.0$ $\mathrm{g} / \mathrm{dL}$, Platelets: $\geqslant 50 \times 10^{\wedge} 9 / \mathrm{L}$, Alanine aminotransferase / Aspartate aminotransferase / Alkaline phosphatase: $\leqslant 5$ times upper limit of normal (ULN), Bilirubin: $\leqslant 3$ times ULN)
- Serum creatinine: within normal limits or $<120 \mu \mathrm{~mol} / \mathrm{L}$ for patients aged 60 years or older.
- $\quad$ Calculated Glomerular filtration rate (GFR) $\geqslant 45 \mathrm{~mL} / \mathrm{min}$.

Exclusion Criteria

- Known hypersensitivity to Gallium-68, to NODAGA, to DOTA, to LM3, or to any of the excipients of Gallium-68 DOTA-LM3 or Gallium-68 NODAGA-LM3.
- Presence of active infection at screening or history of serious infection within the previous 6 weeks.
- Therapeutic use of any somatostatin analog, including long-acting Sandostatin (within 28 days) and short-acting Sandostatin (within 2 days) prior to study imaging. If a patient is on long-acting Sandostatin, then a wash-out phase of 28 days is required before the injection of the study drug. If a patient is on short-acting Sandostatin, then a wash-out phase of 2 days is required before the injection of the study drug.
- Prior or planned administration of a radiopharmaceutical within 8 half-lives of the radionuclide used on such radiopharmaceutical including at any time during the current study.
- Pregnant or breast-feeding women.
- Current history of any malignancy other than neuroendocrine tumor; patients with a secondary tumor in remission of $>5$ years can be included.
- Any mental condition rendering the patient unable to understand the nature, scope and possible consequences of the study, and/or evidence of an uncooperative attitude.

Supplemental Table 2. The SUVmax and tumor-to-background ratios of 38 reference lesions

| Time after injection (minutes) | ${ }^{68} \mathrm{Ga}-$ NODAGA-LM3 $(N=18)$ | ${ }^{68} \mathrm{Ga}$-DOTA-LM3 $(N=20)$ | $P$ value |
| :---: | :---: | :---: | :---: |
|  | SUVmax |  |  |
| 5 | $31.3 \pm 19.7$ | $36.6 \pm 23.6$ | 0.455 |
| 15 | $40.4 \pm 25.9$ | $41.9 \pm 26.5$ | 0.860 |
| 30 | $47.1 \pm 31.0$ | $45.3 \pm 29.3$ | 0.858 |
| 45 | $54.9 \pm 37.3$ | $46.6 \pm 31.2$ | 0.461 |
| 60 | $57.5 \pm 39.4$ | $47.2 \pm 32.6$ | 0.385 |
| 120 | $74.6 \pm 56.3$ | $46.1 \pm 30.9$ | 0.058 |
| Tumor-to-blood-pool ratio |  |  |  |
| 5 | $16.4 \pm 11.7$ | $15.5 \pm 10.0$ | 0.803 |
| 15 | $28.0 \pm 20.1$ | $22.8 \pm 14.7$ | 0.366 |
| 30 | $45.5 \pm 38.6$ | $24.6 \pm 15.8$ | 0.044 |
| 45 | $58.8 \pm 46.4$ | $32.0 \pm 24.4$ | 0.038 |
| 60 | $57.1 \pm 38.9$ | $38.4 \pm 29.1$ | 0.099 |
| 120 | $74.8 \pm 67.2$ | $41.5 \pm 41.4$ | 0.071 |
| Tumor-to-kidney ratio |  |  |  |
| 5 | $1.3 \pm 0.7$ | $4.8 \pm 3.4$ | $<0.001$ |
| 15 | $2.0 \pm 1.1$ | $7.6 \pm 5.9$ | $<0.001$ |
| 30 | $2.4 \pm 1.4$ | $8.4 \pm 6.3$ | $<0.001$ |
| 45 | $3.0 \pm 1.8$ | $10.3 \pm 8.2$ | 0.001 |
| 60 | $3.1 \pm 1.9$ | $11.0 \pm 9.6$ | 0.002 |
| 120 | $4.0 \pm 2.5$ | $10.3 \pm 9.2$ | 0.008 |
| Tumor-to-liver ratio |  |  |  |
| 5 | $4.2 \pm 2.8$ | $12.5 \pm 7.7$ | $<0.001$ |
| 15 | $6.2 \pm 4.3$ | $16.4 \pm 9.2$ | $<0.001$ |
| 30 | $7.8 \pm 5.8$ | $17.6 \pm 9.9$ | 0.001 |
| 45 | $9.4 \pm 7.1$ | $18.6 \pm 11.6$ | 0.006 |
| 60 | $10.0 \pm 7.4$ | $18.5 \pm 11.5$ | 0.011 |
| 120 | $12.6 \pm 9.5$ | $20.7 \pm 12.9$ | 0.035 |

Supplemental Table 3. Residence times and absorbed doses of liver.

|  | Patient \#* | Arm | Residence time (h) | Absorbed doses (mGy/MBq) |
| :---: | :---: | :---: | :---: | :---: |
| Patients with | 1 | A | 0.448 | 0.124 |
| fulminant liver | 2 | B | 0.536 | 0.147 |
| diseases | 7 | B | 0.546 | 0.150 |
|  | 16 | B | 0.719 | 0.196 |
|  | 4 | B | 0.092 | 0.028 |
|  | 5 | A | 0.144 | 0.042 |
| Patients without | 6 | A | 0.172 | 0.050 |
| fulminant liver | 9 | A | 0.160 | 0.046 |
| diseases | 11 | A | 0.165 | 0.048 |
|  | 12 | A | 0.110 | 0.033 |
|  | 13 | B | 0.198 | 0.052 |
|  | 14 | A | 0.169 | 0.049 |
|  | A | 0.181 | 0.052 |  |

* Patient \#3 and \#8 were dropped out.

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