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## PRODUCTION OF HIGH SPECIFIC ACTIVITY Y-86 USING ELECTROCHEMICAL SEPARATION

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**Purpose:** Yttrium-90 is one of the most widely used radionuclides for targeted radiotherapy. However, Y-90 only emits  $\beta$ -particles making accurate dosimetry difficult. Availability of the positron-emitting Y-86 would allow for biodistribution determination and dosimetry calculation on an individual patient basis with PET. The aim of this study is to produce high purity Y-86 in an efficient, cost-effective manner for routine use and supply. **Methods:** Y-86 was produced via the  $^{86}\text{Sr}(\text{p,n})^{86}\text{Y}$  nuclear reaction, 50 mg of enriched  $\text{SrCO}_3$  was irradiated under a 2  $\mu\text{A}$  beam current for <3 hr. The target was dissolved in 2.8M  $\text{HNO}_3$  acid bath. The dissolved solution was transferred to electrochemical cell. The solution was diluted with water, and 1ml of 0.5M  $\text{NH}_4\text{NO}_3$  electrolyte was added. The pH of the solution was adjusted to 2.5-3. The solution was electrolyzed at 1200 mA (40 min) using the two Pt plate-electrodes. A second electrolysis (150 mA for 20 min) was performed in fresh 3 mM  $\text{HNO}_3$  using one Pt plate and the Pt wire as electrodes. The Y-86 was collected from the Pt wire using 2.8M  $\text{HNO}_3/\text{EtOH}$ . After evaporation, Y-86 was reconstituted in 100 $\mu\text{l}$  of 0.1 M HCl. Specific activity was determined via titration of  $^{86}\text{Y}(\text{OAc})_3$  with DOTA. **Results:** Average yields of 2.2 mCi/ $\mu\text{A} \cdot \text{h}$  were achieved which were 58% of theoretical. The major radioisotopic contaminants at EOB were identified to be  $^{86\text{m}}\text{Y}$ ,  $^{87}\text{Y}$ , and  $^{88}\text{Y}$ . Over 95% of the Y-86 was adsorbed on the Pt plate during the first electrolysis, with >97% being re-collected on the Pt wire after the second. **Conclusion:** Y-86 was produced in good yield using a small amount of recyclable  $\text{SrCO}_3$ . The electrochemical cell with three Pt electrodes significantly accelerated the electrodeposition speed of Y-86. High pure Y-86 was reconstituted in a final small volume and DOTA was labeled successfully with purified Y-86.

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## Comparison of in vivo Distribution of Estrogen Receptor $\beta$ Selective [F-<sup>18</sup>]FEDPN in $\alpha/\beta$ ERKO Mice with [F-18]FES

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**Purpose:** Estrogen receptor  $\beta$  (ER $\beta$ ) could be a factor that determines the level of estrogen action in certain estrogen target tissues. ER $\beta$  is found in breast cancer, and its levels relative to ER $\alpha$  decline with disease progression. Thus, the independent quantification of ER $\alpha$  and ER $\beta$  levels in breast cancer by imaging might be predictive of responses to different hormone therapies. **Methods:** The hydroxy group of (2R,3S)-2,3-bis(4-benzyloxyphenyl)-5-hydroxy-pentanitrile was converted to the fluorine compound using DAST and the benzyl groups were removed by hydrogenation to give FEDPN. For the <sup>18</sup>F labeling 5-tosyl-(2R,3S)-2,3-bis(4-methoxyethoxymethyl-phenyl)-pentanenitrile was prepared. This substrate and <sup>18</sup>F were heated 35 sec using a microwave. Following deprotection (3M HCl) and HPLC purification, the <sup>18</sup>F labeled FEDPN was isolated. Biodistribution studies were carried out using immature female Sprague-Dawley rats and ER $\alpha$ - and ER $\beta$ -knockout mice. **Results:** The synthesized FEDPN has an 8.3-fold absolute affinity preference for ER $\beta$ . [<sup>18</sup>F]Fluoride-labeled FEDPN was prepared from a toluenesulfonate precursor, which provided [<sup>18</sup>F]FEDPN with a specific activity greater than 3100 Ci/mmol after HPLC purification. Biodistribution studies revealed specific uptake of [<sup>18</sup>F]FEDPN in the uterus and ovaries. Experiments using ER $\alpha$ - and ER $\beta$ -knockout mice demonstrated the expected ER $\alpha$ -subtype dependence in the tissue uptake of [<sup>18</sup>F]FES, which has a 6,3-fold preference for ER $\alpha$ . The tissue uptake of [<sup>18</sup>F]FEDPN in the ER knockout mice showed some evidence of mediation by ER $\beta$ , but the levels of specific uptake of this agent were relatively modest. **Conclusion:** Based on our results, imaging of ER $\alpha$  can be done effectively with [<sup>18</sup>F]FES, but imaging of ER $\beta$  will likely require agents with more optimized ER $\beta$  binding affinity and selectivity than [<sup>18</sup>F]FEDNP.