### **Activation Products in Proton Therapy**

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# **Topics**

- General considerations
- Determination of activation cross sections
- Beam collimator activation
- Long-lived activation products in biologically relevant elements
- Formation of short-lived  $\beta^+$  emitters in human tissue
- Conclusions

# **General Considerations**

- For therapy purposes, well collimated proton beams are needed
- Proton energy is tailored between 70 and 250 MeV, depending on the application
- In proton therapy atomic and molecular data are of great significance
- Nuclear interactions are of lesser importance, except for cases where high energy secondary particles are emitted
- Data for the formation of activation products are not of paramount importance; however, they are needed in several special contexts

## Determination of Activation Cross Sections in the Medium Energy Range

#### **Experimental Method**



- Irradiation of target material with protons at a low current.
- Very often a stack of thin samples is irradiated (*stacked-foil technique*).
- Calculation of proton energy effective in each sample
- Determination of proton flux via a monitor reaction (or via a Faraday cup)
- Determination of absolute activity of the product nuclide , non-destructively or after chemical separation
- Calculation of cross section
- Construction of excitation function

#### **Irradiation Facilities**

Several types of cyclotrons and accelerators are needed to cover the full energy range.

The Jülich group used following machines:

compact cyclotron ( $\leq$  21 MeV); injector of COSY ( $\leq$  45 MeV); accelerators at PSI ( $\leq$  72 MeV), Uppsala ( $\leq$  180 MeV), iThemba LABS ( $\leq$  200 MeV) and Saclay ( $\leq$  350 MeV)

## **Nuclear Model Calculations**

- Hauser-Feshbach and precompound formalism successfully applied up to 50 MeV
  - Commonly used codes: GNASH, STAPRE, EMPIRE II
- Hybrid exciton model commonly used in higher energy region
  *Common code*: ALICE-IPPE
- Direct interactions needed to be included
- Complex particle emission extremely difficult to treat

## **Activation Cross Section Needs**

- Estimation of collimator activation in proton therapy facilities
- Estimation of long-lived activation products in biologically relevant elements
  - formation of <sup>7</sup>Be
  - formation of <sup>22,24</sup>Na and other medium mass products
- Formation of short-lived  $\beta^+$  emitters in human tissue

# **Activation of Beam Collimators**

- Proton therapy demands high quality beams
- Tailoring of energy and homogenisation of intensity achieved through collimators
- Activation of collimators is of some concern
- Commonly used collimators include titanium, brass, tungsten, etc.

### Results for a Pure Element as Target (Easily detectable products)

#### **Example:** <sup>nat</sup>Cu(p,x)<sup>55,56,58</sup>Co processes



 Model calculations reproduce experimental data well up to E<sub>p</sub> ≤ 120 MeV

## Results for a Pure Element as Target (Difficult to detect product)

#### **Example:** <sup>nat</sup>Ti(p,x)<sup>45</sup>Ca



 Good agreement between experiment and theory over the whole energy range

## **Results for an Alloy as Target**

#### **Example: Formation of** <sup>52,54</sup>Mn from brass



 Model calculation reproduces experimental data with partial success up to proton energies of about 120 MeV.

# **Activation of Brass Collimator**



Proper shielding of therapy facilities is mandatory

# Long-Lived Activation Products in Biologically Relevant Elements

- Biologically relevant elements include H, C, N, O, F, Na, Mg, Si, P, S, Cl, Ca, Fe etc.
- Longer-lived activation products formed during proton therapy may include <sup>7</sup>Be ( $T_{\frac{1}{2}} = 53 \text{ d}$ ), <sup>22</sup>Na ( $T_{\frac{1}{2}} = 2.6 \text{ a}$ ), <sup>24</sup>Na ( $T_{\frac{1}{2}} = 15.0 \text{ h}$ ) and several other medium mass products, like <sup>42</sup>K ( $T_{\frac{1}{2}} = 12.4 \text{ h}$ ), <sup>43</sup>K ( $T_{\frac{1}{2}} = 22.2 \text{ h}$ ), <sup>51</sup>Cr ( $T_{\frac{1}{2}} = 27.7 \text{ d}$ ), <sup>52</sup>Mn ( $T_{\frac{1}{2}} = 5.6 \text{ d}$ ), <sup>54</sup>Mn ( $T_{\frac{1}{2}} = 312 \text{ d}$ ), <sup>55</sup>Co ( $T_{\frac{1}{2}} = 17.5 \text{ h}$ ), etc.
- <sup>7</sup>Be formation in interactions of protons with light elements C, N, O, F and Na involves both <sup>7</sup>Be-emission (as a complex particle) and residual nucleus formation (after emission of several nucleons and α-particles). For heavier target elements, emission of complex particle <sup>7</sup>Be is more probable.
- <sup>22,24</sup>Na and heavier mass radioactive products are formed as residual nuclei

# Systematics of Excitation Functions of (p,<sup>7</sup>Be) Reactions



 Probability of <sup>7</sup>Be emission decreases with increasing mass of the target nucleus

# Formation of <sup>22,24</sup>Na in the Interactions of Protons with <sup>nat</sup>Cl



- Cross sections for the formation of  $^{22,24}$ Na are relatively small (2 5 mb).
- Theory reproduces formation cross section with varying degree of success.

# Formation of Short-lived β<sup>+</sup> Emitters in Human Tissue

Interactions of protons with constituents of human tissue generate short-lived  $\beta^+$  emitters like <sup>11</sup>C (T<sub>1/2</sub> = 20 min), <sup>13</sup>N (T<sub>1/2</sub> = 10 min), <sup>14</sup>O (T<sub>1/2</sub> = 1.15min), <sup>15</sup>O (T<sub>1/2</sub> = 2 min), <sup>18</sup>F (T<sub>1/2</sub> = 110 min), etc.

Examples of contributing nuclear reactions

<sup>12</sup>C(p,pn)<sup>11</sup>C
 <sup>14</sup>N(p,pn)<sup>13</sup>N
 <sup>14</sup>N(p,n)<sup>14</sup>O
 <sup>18</sup>O(p,n)<sup>18</sup>F

 $^{16}O(p,\alpha)^{13}N$  $^{15}N(p,n)^{15}O$ 

 $^{14}N(p,\alpha)^{11}C$ 

<sup>16</sup>O(p,pn)<sup>15</sup>O

#### Significance of production data

 a)Estimation of extra dose due to activation products
 b)PET investigation of the patient after proton therapy (utilising the <sup>11</sup>C formed in the tissue); localises dose distribution in the treated area

# Formation of Short-lived β<sup>+</sup> Emitters

(Protons on human tissue)

#### **Example : <sup>11</sup>C formation**



Improved data base > 50 MeV

# Estimated Activity in Human Tissue and Bone as a Result of Proton Therapy

Assumption: 200 MeV proton, 2 nA, 2 min irradiation

Radionuclide	Activity (MBq)		
	Muscle tissue	Adipose tissue	Bone tissue
<sup>11</sup> C	6.5	19.2	6.4
<sup>13</sup> N	3.9	2.2	2.3
<sup>15</sup> O	2.4	0.7	1.4
Total activity of $\beta^+$ emitters: $10 - 25 \text{ MBq}$ ^7Be activity: $40 \text{ kBq}$ ^22,24 Na activity (in bone): < 250 Bq			

# Conclusions

- Activation of beam collimators at proton therapy facilities is of some concern regarding the therapy personnel.
- Formation of long-lived radioactive products in tissue and bone can be regarded as negligible.
- Formation of short-lived β<sup>+</sup> emitters is of some significance. Total activity (10 – 25 MBq) is sufficient for dose localisation via PET studies; the extra radiation dose from β<sup>+</sup> emitters is, however, negligible (< 1 %).</li>