New cross section measurements for production of the positron emitters ⁷⁵Br and ⁷⁶Br *via* intermediate energy proton induced reactions[†]

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⁷⁵Br / ⁷⁶Br / Intermediate energy / Proton induced reactions / Excitation function / Integral yield

Summary. The radionuclides ⁷⁵Br and ⁷⁶Br are of considerable interest for application in Positron Emission Tomography (PET). For their production so far mostly low-energy or low-yield production processes have been employed. In this work the production of both isotopes was investigated using intermediate energy proton induced reactions and enriched target materials. Cross section data of the ^{77,78,80}Se(p, xn)⁷⁵Br processes were measured up to 66 MeV and those of the ^{77,78,80}Se(p, xn)⁷⁶Br processes up to 85 MeV. The measured excitation functions were compared with calculated results of the nuclear model code ALICE-IPPE. From the experimental excitation curves, thick target yields were calculated. They are fairly high over the investigated intermediate energy ranges. The production possibilities of ⁷⁵Br and ⁷⁶Br *via* various routes and the radionuclidic impurities associated with them are discussed.

1. Introduction

There is a longstanding interest in the radioisotopes of bromine for application in nuclear medicine [1–5]. This is based on their versatile decay properties and covalent chemical bonding. The two radionuclides ⁷⁵Br ($T_{1/2} = 1.6 \, \text{h}$) and ⁷⁶Br ($T_{1/2} = 16.2 \, \text{h}$) are useful for application in positron emission tomography (PET) by virtue of their high positron emission intensities of 75.5% and 57%, respectively. The high nucleophilicity of bromide bears advantages in synthesis of radiolabelled compounds, and n.c.a. products *via* interhalogen-exchange can be obtained easily [2–4]. Further, it can rather easily be oxidised *in situ*, thus facilitating n.c.a. electrophilic substitution reactions in vinyl and aryl positions [5]. The lighter isotope ⁷⁵Br with its short half-life, high positron emission intensity and low positron energy, had been extensively evaluated up to about 1990 for

its potential use in nuclear medicine [6, 7]. A disadvantage associated with this radionuclide, however, is its decay into the long-lived 75 Se ($T_{1/2}=119.8\,\mathrm{d}$) which causes extra radiation dose. On the other hand, the physical properties of 75 Br make this radionuclide interesting for imaging of fast biochemical processes. In recent years, the radionuclide 76 Br has found versatile applications like radiolabeling of antibodies for tumour imaging, or labelling of receptor ligands for brain studies, cf. [8]. The important decay properties of both radionuclides are given in Table 1.

For both radionuclides, and in particular for ⁷⁵Br, a serious drawback is their limited availability. Several routes for the production of 75Br and 76Br had been investigated up to 1986, for reviews cf. [9, 10]. They included formation of both the radionuclides directly as well as via the precursors 75Kr and 76Kr, respectively, obtained through irradiations with protons, deuterons, ³He- and ⁴He-particles. The maximum quantities of 75Br and 76Br achieved were about 4 GBq and 0.5 GBq, respectively. Later reports making use of the reactions 78 Kr $(d, n\alpha)^{75}$ Br [11], 78 Kr $(p, \alpha)^{75}$ Br [12], 74 Se $(d, n)^{75}$ Br [13] and 78 Kr $(d, \alpha)^{76}$ Br [14] led only to limited quantities of about 1 GBq of 75Br and 10 MBq of ⁷⁶Br. In recent years more attention has been paid to the production of ⁷⁶Br, particularly via the ⁷⁶Se(p, n)⁷⁶Br reaction [15, 16]. So far the production of ⁷⁵Br has been successfully done via the 75 As(3 He, 3n) 75 Br and 76 Se(p, 2n) 75 Br reactions, and that of ⁷⁶Br via the ⁷⁵As(³He, 2n)⁷⁶Br and 76 Se $(p, n)^{76}$ Br reactions.

This work is part of a broader study investigating the formation of radiobromine via proton induced reactions on enriched selenium targets and focuses on the production possibilities of the two radionuclides ⁷⁵Br and ⁷⁶Br using intermediate energy protons. Cross section measurements up to 66 MeV were done on the relevant ^{77,78}Se(p, xn)⁷⁵Br and the ^{77,78,80}Se(p, xn)⁷⁶Br processes, using highly enriched target isotopes. In case of the ⁸⁰Se(p, 5n)⁷⁶Br reaction the investigated energy range was extended to 85 MeV. The new excitation functions obtained were compared with results of calculations based on the precompound nuclear model code ALICE-IPPE. The integral yields for the production of the two radionuclides were also calculated, and a discussion of the results with special respect to radionuclidic purity

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536 I. Spahn et al.

Table 1. Decay properties of the radionuclides ⁷⁵Br and ⁷⁶Br and their investigated production routes.

	Half- life (h)	Decay mode (%)	$ \beta^{+}_{\text{max}} $ energy (MeV)	Main γ-rays in keV (%)	Previously investigated production routes	Q-value (MeV)
⁷⁵ Br	1.6	β ⁺ (75.5) EC (24.5)	1.72	141.3 (6.6) 286.5 (88)	$^{78}{ m Kr}(d,nlpha)$ $^{78}{ m Kr}(p,lpha)$ $^{75}{ m As}(^3{ m He},3n)$ $^{76}{ m Se}(p,2n)$	-2.041 -0.177 -13.176 -14.967 -1.991
⁷⁶ Br	16.2	β ⁺ (58.2) EC (41.8)	3.94	559.1 (74) 657.0 (15.9)	75 As(3 He, 2 n) 76 Se(2 n, 1 n) 79 Br(2 n, 4 n) 76 Kr \rightarrow 76 Br 81 Br(2 n, 76 Kr \rightarrow 76 Br	-3.954 -5.745 -32.051 -52.324

Decay data taken from [27, 28, 29 (β ⁺ intensity)].

is given. The formation of other radioisotopes of bromine, namely ⁷⁷Br and ^{80m}Br, and their implication on radionuclidic purity will be discussed in a subsequent paper.

2. Experimental

Cross sections were measured by activation and identification of the radioactive products. The relevant techniques are given below.

2.1 Sample preparation

For investigation of different nuclear processes, targets of enriched elemental ⁷⁷Se, ⁷⁸Se and ⁸⁰Se powder were used, the detailed compositions of which are given in Table 2. In order to obtain thin samples for irradiations, the sedimentation technique described in detail earlier [16, 17] was used. The powder was sedimented from ethanol on a 0.1 mm thick aluminium backing and covered with 0.01 mm thick Al foil for protection. This way sediment layers could be obtained, each with enriched Se isotope content of 2.9 to 28.9 mg/cm². The diameter of the deposited layer was either 7 or 10 mm.

2.2 Irradiations and beam current measurement

The samples were irradiated in a stacked-foil arrangement together with different monitor foils. The proton energy degradation within the stack was calculated at Jülich with the computer code STACK which is based on the Bethe formalism and the tables of Williamson *et al.* [18] and with a similar code at iThemba LABS in Somerset West.

Table 2. Isotopic compositions of used enriched Se target material.

	⁷⁶ Se (%)	⁷⁷ Se (%)	⁷⁸ Se (%)	⁸⁰ Se (%)	⁸² Se (%)
⁷⁷ Se ^a ⁷⁸ Se ^a ⁸⁰ Se ^b	1.21	91.77	4.68	1.87	0.41
	0.11	0.17	98.58	1.00	0.08
	0.01	< 0.005	0.008	99.90	0.07

a: Material provided by Oak Ridge National Laboratory;

Irradiations up to 45 MeV proton energy were done at the injector cyclotron of COSY of the Forschungszentrum Jülich, Germany. Irradiations in the higher energy region were carried out at the Separate Sector Cyclotron (SSC) of iThemba LABS in Somerset West, South Africa. The proton energy range covered by the irradiations was from 21 to 80 MeV. The thin ⁷⁷Se and ⁷⁸Se targets were irradiated at both cyclotrons with protons of energies up to 66 MeV. The energy ranges covered in the stacks were chosen such that they partly overlapped, thereby allowing a comparison of results in the overlapping regions, obtained in independent measurements.

The beam current was measured *via* current integration on one hand, and *via* monitor reactions, on the other. As monitors thin Al, Cu and Ni foils were used, inducing the corresponding 27 Al $(p, x)^{22}$ Na, nat Cu $(p, xn)^{62,63}$ Zn and nat Ni $(p, x)^{57}$ Ni processes, the cross sections of which are well known [19]. At iThemba LABS current measurement was also done using the Faraday cup described in detail earlier [20]. The two methods of beam current determination showed an agreement within about 5%.

2.3 Measurement of radioactivity

The induced radioactivity was measured using HPGe detector γ -ray spectrometry, analysing all major γ -rays of the investigated radionuclides. Several spectra, registered at various times after the end of bombardment (EOB), were analysed and the results averaged. The peak area analysis was done using either the software Gamma Vision 6.01 (EG&G Ortec) or Emca 2000 2.03.0 (Silena Int. spa). All γ -ray measurements were done non-destructively. Detectors were calibrated and their efficiencies determined using standard sources from PTB Braunschweig and Amersham International. All measurements were done at a minimum distance of between 10 and 50 cm to the detector to avoid coincidence losses.

For the study of 75 Br the two strongest γ -rays of energies 141.32 and 286.57 keV were analysed, and for 76 Br the 559.10 and 657.36 keV γ -rays were considered. Since both γ -rays are also emitted in the decay of 76 As, a small contribution of that nuclide in 76 Br cannot be ruled out.

b: Material provided by Chemotrade GmbH.

However, this contribution is expected to be small because the cross sections for the nuclear processes 77 Se $(p, 2p)^{76}$ As, 78 Se $(p, ^3\text{He})^{76}$ As and 80 Se $(p, \alpha n)^{76}$ As are expected to be rather small, in comparison to the respective (p, xn) reactions leading to the formation of 76 Br. To verify this, decay curve analyses were done to check for any deviation in the measured half-life due to contribution of the longer lived 76 As $(T_{1/2}=26.4\,\text{h})$. The analyses confirmed the literature value with an average uncertainty of 7%, which is well within the experimental uncertainties and can be considered as a maximum possible contamination.

2.4 Calculation of reaction cross sections and uncertainties

Cross sections were calculated using the well-known activation equation. As mentioned above, results obtained from the analysis of several clear γ -lines belonging to the same nuclide were averaged. The enriched isotopes had different abundances. However, all cross section values were normalised to 100% enrichment of the target isotope.

The total uncertainties of the measurements were calculated with the error propagation according to Gauss. They are mainly based on the individual uncertainties concerning the incident proton energies, determination of particle flux and detector efficiency. In general, the total uncertainty in cross section amounted to between 9 and 17%.

3. Results and discussion

3.1 Reaction cross sections

The measured nuclear reaction cross sections relevant to the production of ⁷⁵Br and ⁷⁶Br are given in Tables 3 and 4, respectively, together with their corresponding uncertainties. The results of the accomplished experiments, done at two cyclotrons in different facilities, showed good consistency. Regarding the literature data, the formation of ⁷⁶Br up to 35 MeV was determined well by Hassan et al. [16]. Concerning the production of the radionuclide ⁷⁵Br, results have been reported for proton induced reactions on nat Se and ⁷⁶Se up to about 40 MeV [16, 21, 22]. A few other data are also available up to proton energies of 30 MeV [23]. All those data are discussed below. In the higher proton energy range, to date our measurements represent the first (p, xn)reaction cross section data for the production of these radionuclides. Based on those experimental values, obtained using enriched target material, excitation functions could be constructed as shown in Figs. 1–5. For comparison, existing experimental results in the literature are given, where this was possible.

Formation of 75Br

Fig. 1 depicts the cross section data for the production of 75 Br via the 77 Se(p, 3n) process. The excitation curve shows a maximum of about 250 mb at 34 MeV. The previously published data of Levkovskij [23] are rather high. A recent publication by Takács *et al.* [24] suggests a general downward adjustment of those data up to 20%, which is shown in the figure and leads to better agreement with our meas-

Table 3. Reaction cross sections in mb for the formation of 75Br.

Energy	75 Bi	r
(MeV)	$^{77}\mathrm{Se}(p,3n)$	78 Se $(p,4n)$
24.3 ± 1.0	27.4 ± 3.6	
26.6 ± 0.8	26.3 ± 3.5	
28.7 ± 0.7	106.7 ± 13.9	
31.1 ± 0.5	178.4 ± 23.2	
31.5 ± 1.0	182.6 ± 23.7	
32.2 ± 0.9		0.5 ± 0.1
32.5 ± 0.6		1.1 ± 0.1
34.2 ± 0.4		0.6 ± 0.1
35.6 ± 0.4		6.9 ± 0.9
35.7 ± 0.4	267.5 ± 34.8	
35.8 ± 0.6		0.4 ± 0.0
37.0 ± 0.4	161.2 ± 21.0	
37.2 ± 0.3		6.3 ± 0.8
38.4 ± 0.3		11.9 ± 1.6
38.6 ± 0.3	188.2 ± 24.5	
39.9 ± 0.3	101.1 ± 13.1	
39.9 ± 0.2		29.5 ± 3.8
39.9 ± 0.5	116.4 ± 104.8	
40.8 ± 0.2		35.7 ± 4.6
41.4 ± 0.2	97.8 ± 12.7	
41.8 ± 0.2		43.1 ± 5.6
42.4 ± 0.2		35.6 ± 4.6
43.2 ± 0.2		53.9 ± 7.0
43.2 ± 0.5		36.3 ± 3.9
43.7 ± 0.2	63.9 ± 8.3	
44.1 ± 0.2		71.6 ± 9.3
44.5 ± 0.2	87.3 ± 11.4	
44.9 ± 0.2		75.8 ± 9.8
48.4 ± 0.5	37.4 ± 6.4	
53.6 ± 0.4		57.6 ± 6.2
56.6 ± 0.3	28.3 ± 4.8	
59.3 ± 0.3		42.8 ± 4.6
64.7 ± 0.2		31.7 ± 3.0

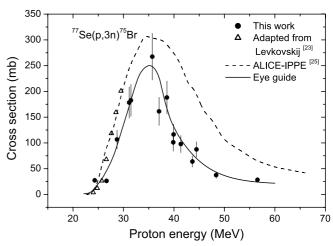


Fig. 1. Experimentally determined excitation function of the 77 Se(p, 3n) 75 Br reaction together with data from the literature and results of ALICE-IPPE calculation.

urements. The remaining shift in energy may be within the limits of uncertainty, which is, however, not given in the case of the Levkovskij data which were taken from the EXFOR file [28]. The dashed line in Fig. 1 describes theoretical results obtained using the ALICE-IPPE code, as given in the MENDL-2P data file [25]. Evidently, the model calculation reproduces the shape of the excitation function well, but generally overestimates the cross section data.

538 I. Spahn et al.

Table 4. Reaction cross sections in mb for the formation of ⁷⁶Br.

Energy (MeV)	$^{77}\mathrm{Se}(p,2n)$	78 Se $(p,3n)$	${}^{80}\mathrm{Se}(p,5n)$
21.3±0.9	478.5 ± 43.1		
23.4 ± 0.8		1.0 ± 0.1	
24.3 ± 0.8	446.0 ± 40.1	10.5 1.6	
25.2 ± 0.6 26.6 ± 0.8	332.7 ± 29.9	12.5 ± 1.6	
26.8 ± 0.8	332.1 ± 29.9	5.6 ± 0.5	
26.9 ± 0.5		30.2 ± 3.9	
27.8 ± 0.5		133.3 ± 17.3	
28.7 ± 0.5	157.6 ± 20.5		
29.5 ± 0.5		81.3 ± 10.6	
29.9 ± 0.5 30.9 ± 0.7		176.2 ± 22.9 66.2 ± 6.0	
31.1 ± 0.5	110.1 ± 14.3	00.2 ± 0.0	
31.5 ± 0.5	122.0 ± 15.9		
32.2 ± 0.4		233.3 ± 30.3	
32.5 ± 0.4	00.2 10.0	207.5 ± 27.0	
33.7 ± 0.4 34.2 ± 0.4	99.2 ± 10.9 64.8 ± 7.1	212.4 ± 40.6	
34.2 ± 0.4 35.6 ± 0.4	04.6 ± 7.1	312.4 ± 40.6 267.2 ± 34.7	
35.7 ± 0.4	87.4 ± 11.4	207.2 ± 3 1.7	
37.0 ± 0.3	66.0 ± 8.6		
37.2 ± 0.3		323.0 ± 42.0	
38.4 ± 0.6 38.6 ± 0.3	91.5 ± 10.6	222.1 ± 28.9	
38.0 ± 0.3 39.9 ± 0.2	81.5 ± 10.6 49.1 ± 6.4		
39.9 ± 0.2	47.1 ± 0.4	265.3 ± 34.5	
39.9 ± 0.5	70.8 ± 6.4		
40.8 ± 0.2		216.8 ± 28.2	
41.4 ± 0.2	47.8 ± 6.2	100 0 1 26 0	
41.8 ± 0.2 42.4 ± 0.2		199.9 ± 26.0 177.7 ± 23.1	
43.2 ± 0.2		164.4 ± 21.4	
43.2 ± 0.5		188.8 ± 17.0	
43.7 ± 0.2	30.9 ± 4.0		
44.1 ± 0.2	26.1 4.7	121.3 ± 15.8	
44.5 ± 0.2 44.9 ± 0.2	36.1 ± 4.7	155.8 ± 20.3	
45.0 ± 0.2		131.4 ± 11.8	
46.2 ± 0.4			5.4 ± 0.5
48.4 ± 0.3	37.2 ± 3.3		
48.8 ± 0.3			13.0 ± 1.2
51.3 ± 0.3 53.6 ± 0.4		57.9 ± 5.2	20.3 ± 1.8
53.0 ± 0.4 53.9 ± 0.3		31.9 ± 3.2	49.0 ± 4.4
56.5 ± 0.3			56.9 ± 5.1
56.6 ± 0.2	34.6 ± 3.1		
58.9 ± 0.2		~ 4 0 1 4 5	49.2 ± 4.4
59.3 ± 0.3 61.4 ± 0.2		51.0 ± 4.6	73.1 ± 6.6
61.4 ± 0.2 64.7 ± 0.2		46.7 ± 5.1	73.1±0.0
64.7 ± 0.2		46.7 ± 4.2	
66.5 ± 0.4			73.3 ± 6.6
68.5 ± 0.4			50.9 ± 4.6
70.5 ± 0.4 72.4 ± 0.3			55.9 ± 5.0
72.4 ± 0.3 74.5 ± 0.3			53.2 ± 4.8 54.9 ± 4.9
76.5 ± 0.3			45.4 ± 4.1
78.5 ± 0.3			28.4 ± 2.6
80.4 ± 0.2			41.3 ± 3.7
82.3 ± 0.2 84.2 ± 0.2			16.0 ± 1.4 22.0 ± 2.0
$v_{T,2} \perp v_{*2}$			22.0 ± 2.0

The experimental results on the formation of 75 Br via the 78 Se(p, 4n) process are shown in Fig. 2. Those cross sections have been measured for the first time using enriched 78 Se as target material. The suggested excitation function (Fig. 2)

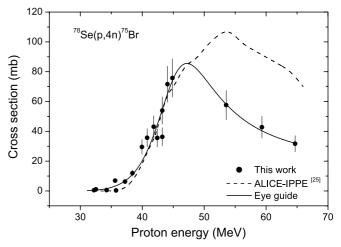


Fig. 2. Experimentally determined excitation function of the ⁷⁸Se(p, 4n) ⁷⁵Br reaction together with the results of ALICE-IPPE calculation.

shows a maximum value of 80 mb at about 47 MeV proton energy. However, additional experimental work is required to define clearly the cross section maximum. The results of the ALICE-IPPE calculation agree very well with the experimental data up to about 45 MeV, but differ considerably thereafter with regard to both shape and magnitude of the excitation function. The comparison points out the difficulties in modeling multi-particle emission from medium mass target nuclei.

Formation of ⁷⁶Br

Figs. 3–5 show the measured cross section values for the formation of 76 Br via three reaction channels, namely (p, 2n), (p, 3n) and (p, 5n), on enriched selenium targets. The existing literature data on the 77 Se $(p, 2n)^{76}$ Br reaction given by Hassan $et\ al.$ [16] and Levkovskij [23] are also shown in Fig. 3. The experimental results obtained in this work are not sufficient to indicate a maximum of the excitation function. The literature shows significantly larger values below 30 MeV. However, the data of Levkovskij, decreased by 20% as suggested by Takács $et\ al.$ [24], come nearer to our measurements. Similarly, the high maximum cross

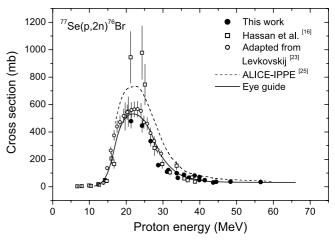


Fig. 3. Excitation function of the 77 Se $(p, 2n)^{76}$ Br reaction compared with data from the literature and results of ALICE-IPPE calculation.

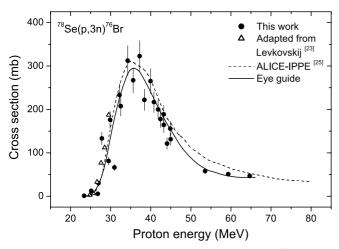


Fig. 4. Experimentally determined excitation function of the 78 Se(p, 3n) 76 Br reaction compared with data from the literature and results of ALICE-IPPE calculation.

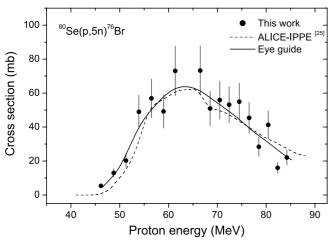


Fig. 5. Excitation function of the 80 Se $(p,5n)^{76}$ Br reaction together with results of ALICE-IPPE calculation.

section of about 900 mb, reported by Hassan *et al.* [16], is based on only 3 points, and may be in error. Our results reported in this work were obtained through three independent experiments in two different laboratories. Therefore there is more confidence in the present values. Thus the eye guide curve shown in Fig. 3 is based on the present data and the adapted values of Levkovskij. As regards the model calculation, again it reproduces the shape of the excitation function well but the magnitude of cross sections is rather high.

The excitation function of the 78 Se(p, 3n) 76 Br reaction is shown in Fig. 4 together with data from the literature and the results of the ALICE-IPPE calculation taken from the MENDL-2P data file [25]. Although the data show some scatter, a clear maximum can be distinguished at 36 MeV. Within the limits of uncertainty the literature data agree well with our results. The agreement between experimental and calculated data is also rather good.

The excitation function of the ${}^{80}Se(p, 5n)^{76}Br$ reaction is shown in Fig. 5. No data were found on this nuclear reaction in the literature but the calculated results from the MENDL-2P data file were available and are shown in the figure. The theoretical curve agrees well with the experimental results,

which, however, show rather large scatter. Additional measurements in the intermediate energy range are necessary to define this curve more precisely.

3.2 Calculated integral yields

Based on the excitation functions determined in this work, the possible maximum production yields of the two radionuclides 75Br and 76Br were calculated using the whole investigated energy range. The results are given in Table 5 for the respective Se isotope used as target material. For comparison the calculated yields based on the irradiation of enriched ⁷⁶Se and reported in the literature [26] are also given in Table 5. Evidently, the yield of 76 Br *via* the (p, 2n)and (p, 3n) processes is considerably higher than that via the (p, n) reaction on ⁷⁶Se. The production of ⁷⁵Br *via* the 77 Se(p, 3n) and 78 Se(p, 4n) processes proves to be less effective than the application of the 76 Se $(p, 2n)^{75}$ Br reaction regarding the investigated energy range. Based on the determined excitation functions higher incident proton energies may, however, further increase the yields of the (p, 3n) and (p, 4n) reactions. An estimation of the respective theoretical yields based on the cross section data given by the ALICE-IPPE code [25] supports this conclusion.

Table 5. Calculated total thick target yields of the produced $^{75}\mathrm{Br}$ and $^{76}\mathrm{Br}$.

Radionuclide	Target nuclide	Energy range [MeV]	Prod. yield [MBq/(μAh)]
⁷⁵ Br ^a	⁷⁷ Se ⁷⁸ Se	$62 \rightarrow 20$ $66 \rightarrow 22$	6153 3350
⁷⁶ Br ^b	⁷⁷ Se ⁷⁸ Se ⁸⁰ Se	$66 \rightarrow 14$ $66 \rightarrow 24$ $86 \rightarrow 34$	1320 1118 539

- b: The calculated yield of ⁷⁶Br *via* the ⁷⁶Se(p, n)-reaction over $E_p = 15 \rightarrow 8$ MeV amounts to 360 MBg/ μ A h [26].

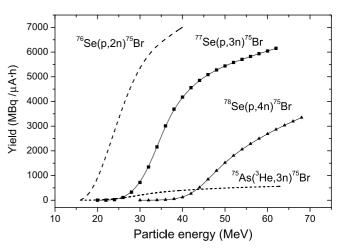


Fig. 6. Integral yields of ⁷⁵Br in proton induced reactions on enriched ⁷⁷Se and ⁷⁸Se calculated from the excitation functions measured in this work. The data for the reactions 76 Se $(p, 2n)^{75}$ Br [16, 22] and 75 As $(^{3}$ He, $^{3}n)^{75}$ Br [30] were taken from the literature.

In Fig. 6 the calculated thick target yields of 75 Br from all investigated nuclear reaction channels are shown as function of particle energy. Obviously, in the energy range below 30 MeV the 76 Se(p, 2n) 75 Br reaction discussed above is very useful for production. The calculated yield of the 75 As(3 He, 3n) 75 Br process [30] is rather low, but this reaction was used for 75 Br production due to the advantageous development of high-current As-containing target material. The calculated thick target yields of 76 Br are shown in Fig. 7 in comparison with the results of the alternative production routes utilizing the 76 Se(p, n) 76 Br and 75 As(3 He, 2n) 76 Br reactions. The yield of the latter is much lower than those of the investigated proton induced reactions.

Concerning the isotopic purity in the production of ⁷⁶Br with intermediate energy protons, the co-produced ⁷⁵Br is the most significant radionuclidic impurity. However, its short half-life allows for a nearly complete decay of this impurity within one half-life of the ⁷⁶Br ($T_{1/2} = 16.2 \,\mathrm{h}$). The production of radionuclidically pure ⁷⁵Br on the other hand, is not possible. In Fig. 8 the production yields of ⁷⁵Br *via* the ⁷⁷Se(p, 3n)⁷⁵Br and ⁷⁸Se(p, 4n)⁷⁵Br reactions, investigated in this work, are shown again, this time together with the corresponding fraction of co-produced

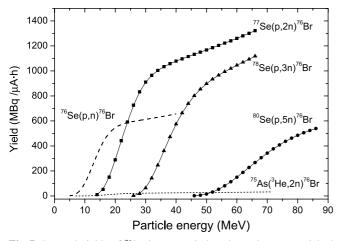


Fig. 7. Integral yields of ⁷⁶Br in proton induced reactions on enriched ⁷⁷Se, ⁷⁸Se and ⁸⁰Se calculated from the excitation functions measured in this work. The data for the reactions ⁷⁶Se(p, n)⁷⁶Br [16, 22] and ⁷⁵As(³He, 2n)⁷⁶Br [30] were taken from the literature.

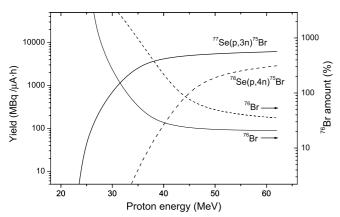


Fig. 8. Calculated thick target yields of 75 Br via the 77 Se(p, 3n) and 78 Se(p, 4n) processes, together with the corresponding ratio of the 76 Br impurity (in %).

⁷⁶Br. The irradiation of a thick ⁷⁷Se target with 66 MeV protons results in an impurity level of about 21% ⁷⁶Br. With regard to the bombardment of ⁷⁸Se, this ratio increases to about 35%. The co-production of the radionuclide ⁷⁷Br and its effect on the isotopic purity will be discussed in detail in a separate publication, as it is subject to several factors like proton energy and irradiation time.

It can be concluded that the application of protons in the intermediate energy range is advantageous for the production of 76 Br. With regard to the production of 75 Br, the intermediate energy reactions are inferior to the production *via* the (p, 2n) process on enriched 76 Se [26], which is due to the lower radionuclide yield as well as the low radionuclidic purity connected to it.

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References

- Stöcklin, G.: Bromine-77 and iodine-123 radiopharmaceuticals. Int. J. Appl. Radiat. Isot. 28, 131–147 (1977).
- Coenen, H. H., Moerlein, S. M., Stöcklin, G.: No-carrier-added radiohalogenation methods with heavy halogens. Radiochim. Acta 34, 47–68 (1983).
- 3. Maziere, B., Loc'h, C.: Radiopharmaceuticals labelled with bromine isotopes. Appl. Radiat. Isot. 37, 703–713 (1986).
- Stöcklin, G.: Molecules labelled with positron emitting halogens. Nucl. Med. Biol. 13, 109–118 (1986).
- Coenen, H. H.: New radiohalogenation methods: An overview. In: *Progress in Radiopharmacology*. (Cox, P. H., Mather, S. J., Sampson, C. B., Lazarus, C. R., eds.) Martinus Nijhoff Publishers, Dordrecht (1986), pp. 196–220.
- Glaser, M., Luthra, S. K., Brady, F.: Applications of positronemitting halogens in PET oncology (Review). Int. J. Oncol. 22, 253–267 (2003).
- Qaim, S. M., Clark, J. C., Crouzel, C., Guillaume, M., Helmeke, H. J., Nebeling, B., Pike, V. W., Stöcklin, G.: PET radionuclide production. In: *Radiopharmaceuticals for Positron Emission Tomography*. (Stöcklin, G., Pike, V. W., eds.) Kluwer Academic Publishers, Dordrecht (1993), pp. 1–43.
- Rowland, D. J., McCarthy, T. J., Welch, M. J.: Radiobromine for imaging and therapy. In: *Handbook of Radiopharmaceuticals*. (Welch, M. J., Redvanly, C. S., eds.) John Wiley & Sons, New York (2003), pp. 441–465.
- Qaim, S. M., Stöcklin, G.: Production of some medically important short-lived neutron-deficient radioisotopes of halogens. Radiochim. Acta 34, 25–40 (1983).
- Qaim, S. M.: Recent developments in the production of ¹⁸F, ^{75,76,77}Br and ¹²³I. Appl. Radiat. Isot. 37, 803–810 (1986).
- 11. De Jesus, O. T., Friedman, A. M.: Production of ⁷⁵Br *via* the 78 Kr $(d, n\alpha)$ ⁷⁵Br reaction. Appl. Radiat. Isot. **39**, 709–710 (1988).
- 12. Tárkányi, F., Kovács, Z., Qaim, S. M.: Excitation functions of proton induced nuclear reactions on highly enriched ⁷⁸Kr: relevance to the production of ⁷⁵Br and ⁷⁷Br at a small cyclotron. Appl. Radiat. Isot. 44, 1105–1111 (1993).
- 13. Qaim, S. M., Stöcklin, G.: Excitation functions of ⁷⁴Se(*d*, *xn*) ^{75,74m}Br reactions: Comparative evaluation of possible routes for the production of ⁷⁵Br at a small cyclotron. Appl. Radiat. Isot. **44**, 1443–1447 (1993).
- 14. Scholten, B., Takács, S., Tárkányi, F., Coenen, H. H., Qaim, S. M.: Excitation functions of deuteron induced nuclear reactions on enriched ⁷⁸Kr with particular relevance to the production of ⁷⁶Br. Radiochim. Acta **92**, 203–207 (2004).

- Tolmachev, V., Lovquist, A., Einarsson, L., Schultz, J., Lundquist, H.: Production of ⁷⁶Br by a low-energy cyclotron. Appl. Radiat. Isot. 49, 1537–1540 (1998).
- 16. Hassan, H. E., Qaim, S. M., Shubin, Yu., Azzam, A., Morsy, M., Coenen, H. H.: Experimental studies and nuclear model calculations on proton-induced reactions on nat Se, 76Se and 77Se with particular reference to the production of of the medically interesting radionuclides 76Br and 77Br. Appl. Radiat. Isot. 60, 899–909 (2004)
- 17. Spahn, I., Steyn, G. F., Nortier, F. M., Coenen, H. H., Qaim, S. M.: Excitation functions of $^{\text{nat}}\text{Ge}(p, xn)^{71,72,73,74}\text{As}$ reactions up to 100 MeV with a focus on the production of ^{72}As for medical and ^{73}As for environmental studies. Appl. Radiat. Isot. **65**, 1057–1064 (2007).
- 18. Williamson, C. F., Boujot, J. P., Picard, J.: Tables of range and stopping power of chemical elements for charged particles of energies 0.5 to 500 MeV. Rapport CEA-R 3042, Saclay, France (1996).
- Tárkányi, F., Takács, S., Gul, K., Hermanne, A. Mustafa, M. G., Nortier, M., Obložinský, P., Qaim, S. M., Scholten, B., Shubin, Yu. N., Zhuang, Y.: Monitor reactions. In: Charged particle cross section database for medical radioisotope production. IAEA-TECDOC-1211, International Atomic Energy Agency, Vienna (2001), pp. 49–152.
- Szelecsényi, F., Steyn, G. F., Kovács, Z., Vermeulen, C., van der Meulen, N. P., Dolley, S. G., van der Walt, N. T., Suzuki, K., Mukai, K.: Investigation of the ⁶⁶Zn(p, 2n)⁶⁴Cu and the ⁶⁸Zn(p, x) ⁶⁴Cu nuclear processes up to 100 MeV: Production of ⁶⁴Cu. Nucl. Instrum. Methods Phys. Res. B **240**, 625–637 (2005).
- Paans, A. M. J., Welleweerd, J., Vaalburg, W., Reiffers, S., Woldring, M. G.: Excitation functions for the production of bromine-75: A potential nuclide for the labelling of radiopharmaceuticals. Appl. Radiat. Isot. 31, 267–273 (1980).

- 22. Kovács, Z., Blessing, G., Qaim, S. M., Stöcklin, G.: Production of ⁷⁵Br *via* the ⁷⁶Se(*p*, 2*n*)⁷⁵Br reaction at a compact cyclotron. Appl. Radiat. Isot. **36**, 635–642 (1985).
- 23. Levkovskij, V. N.: Activation Cross Section Nuclides of Average Masses (A = 40–100) by Protons and Alpha-Particles with Average Energies (E = 10–50 MeV). Inter Vesi, Moscow (1991).
- 24. Takács, S., Tárkányi, F, Sonck, M., Hermanne, A.: Investigation of the ^{nat}Mo(p, x)^{96mg}Tc nuclear reaction to monitor proton beams: New measurements and consequences on the earlier reported data. Nucl. Instrum. Methods Phys. Res. B **198**, 183–196 (2002).
- Shubin, Yu. N., Lunev, V. P., Konobeyev, A. Yu., Dityuk, A. I.: Proton reaction data library for nuclear activation, MENDL-2P (Medium Energy Nuclear Data Library). IAEA-NDS-204, International Atomic Energy Agency, Vienna (1998).
- Qaim, S. M.: Decay data and production yields of some nonstandard positron emitters used in PET. Q. J. Nucl. Med. Mol. Imaging 52, 111–120 (2008).
- 27. Firestone, R. B.: *Table of Isotopes*. Cd ROM edition, Version 1.0. Wiley Interscience, New York (1996).
- 28. Experimental Nuclear Reaction Data (EXFOR), Database Version of June 4 (2008), Nuclear Data Section, IAEA (http://www-nds.iaea.org/exfor/exfor00.htm).
- Qaim, S. M., Bisinger, T., Hilgers, K., Nayak, D., Coenen, H. H.: Positron emission intensities in the decay of ⁶⁴Cu, ⁷⁶Br and ¹²⁴I. Radiochim. Acta 95, 67–73 (2007).
- 30. Alfassi, Z. B., Weinreich, R.: The production of positron emitters 75 Br and 76 Br: excitation functions and yields for 3 He and α -particle induced reactions on arsenic. Radiochim. Acta **30**, 67–71 (1982).
- 31. Misaelides, P., Krauskopf, J., Wolf, G. Bethge, K.: Determination of carbon and oxygen in GaAs by means of ³He-activation. Nucl. Instrum. Methods B **18**, 281–285 (1986).