

Review

How to produce high specific activity tin-117 m using alpha particle beam

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HIGHLIGHTS

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ABSTRACT

Tin-117m is an interesting radionuclide for both diagnosis and therapy, thanks to the gamma-ray and electron emissions, respectively, resulting from its decay to tin-117g. The high specific activity of tin-117 m is required in many medical applications, and it can be obtained using a high energy alpha particle beam and a cadmium target. The experiments performed at the ARRONAX cyclotron (Nantes, France) using an alpha particle beam delivered at 67.4 MeV provide a measurement of the excitation function of the Cd-nat(α,x)Sn-117 m reaction and the produced contaminants. The Cd-116($\alpha,3n$)Sn-117m production cross section has been deduced from these experimental results using natural cadmium. Both production yield and specific activity as a function of the projectile energy have been calculated. These informations help to optimize the irradiation conditions to produce tin-117 m with the required specific activity using α particles with a cadmium target.

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1. Introduction

1.1. The medical interest of tin-117 m

It has been demonstrated since long ago, that β^- emitters are quite effective to relieve bone metastasis pain; in the 60 s for phosphorus-32 (Joshi et al., 1965) and in the 90 s for strontium-89 (Porter et al., 1993), samarium-153 (Collins et al., 1993; Resche et al., 1997) and rhenium-186 g (Maxon et al., 1991; De Klerk et al., 1997). For these β^- emitters, the mean energy (maximum energy) of the emitted electrons varies from one isotope to another, leading to a range in tissues from 0.4 mm to 3 mm (from 2 mm to 8 mm). Since the emitted electrons follow a certain energy distribution, even if these β^- emitters show a good vectorization to the bone metastases, there are always electrons irradiating the radiosensitive bone marrow (Srivastava et al., 1998). This can be avoided by using fixed energy electrons from electron conversion emitters. It is then possible, by choosing the right radionuclide, to have a better control of the dose distribution, with a higher dose on painful bone metastases and a lower dose in the bone marrow. An article from 1973 (Yano et al., 1973) presents the interest of tin-117 m in medicine from the study of several chelates in rats. The study shows the high uptake of tin-117 m in the bones, with a low absorption in the normal tissues. Tin-117 m is a conversion electron emitter (129.36 keV (65.7%), 129.36 keV (11.65%) and 151.56 keV (26.5%)) with a half-life of 13.60 (4) days (Kinsey et al., 2016; Ekström and Firestone, 2004). The emitted electrons have a range in water from 0.2 to 0.3 mm, depending on the considered electron. In 1998, results on dosimetric calculations were published, showing a comparison between the dose received by the bones, and that received by the bone marrow in the case of β^- emitters and Sn-117 m (Srivastava et al., 1998). The results showed that the dose ratio bone/bone marrow is, for Sn-117 m, 4.1 times higher than that of Sr-89, which in turn is 2.7 times higher than that of Sm-153 and 1.2 times higher than that of Re-186. The therapeutic advantage of the conversion electrons emitted by tin-117 m, compared with β^- emitters, has been experimentally shown in 2000 (Bishayee et al., 2000). In addition to conversion electrons, gamma emissions also occur during Sn-117 m decay. In particular, it emits a 158 keV gamma ray (branching ratio 87%), which is suitable for SPECT imaging. After decay, the Sn-117 m changes into the stable Sn-117 g nuclide. Sn-117 m can then be used as a theranostic agent (Srivastava, 2014) thanks to the gamma and conversion electron emission. The Clear Vascular Company in Texas (USA) is dedicated to clinical trials with special emphasis on the diagnosis and therapy of vulnerable plaques (Clear Vascular Inc., 2015; Lafont, 2003) and other inflammatory diseases. This company has developed an injectable radiopharmaceutical called “Tin-

Annexin”, composed by tin-117 m chelated to Annexin V using a DOTA chelate. Pre-clinical and clinical trials, using this radiopharmaceutical made of Annexin V labelled with Sn-117 m, have led to the diagnosis of vulnerable plaques, showing its therapeutic effects on them (Clear Vascular Inc., 2015).

1.2. The tin-117 m production routes

Tin-117 m can be produced by the neutron activation of tin-116 or tin-117 g in reactors. However, its production requires between 2 and 3 weeks of irradiation, and the final product has a low specific activity of the order of the Ci/g. A specific activity between 100 and 1000 Ci/g could be obtained with the use of an additional step of electromagnetic mass separation (Clear Vascular Inc., 2015). Another way to get high specific activity product is to use charged particles (protons, deuterons or α particles) as projectiles. Two main production routes have been identified for the tin-117 m production. The Sb-nat(p,x)Sn-117 m reaction has been studied by the Institute for Nuclear Research (Russian Academy of Science, 2015). Data have been published in 2007 by Ermolaev et al., (2007) and in 2013 by Takács et al. (2013). With this reaction, the expected specific activity is close to 1 kCi/g. Another way is to use α particles as projectiles impinging on a Cd-116 enriched target. Even if there is a limited number of accelerators delivering high energy and high intensity α particle beams, this route is currently used to produce tin-117 m for clinical trials. Indeed, due to its half-life of several days, the irradiation can be performed in few facilities whereas the target is sent to a radiopharmaceutical company, before its deliver to treatment centers. Two data sets have been published for this reaction up to 42 MeV. Therefore, there is a lack of data at high energy, which is unfortunate since industrial production uses energies above 42 MeV. One thick target production yield estimation has been made on the energy range of 47–20 MeV, based on Qaim and Döhler (1984) calculations, and leads to 150 μ Ci/ μ A h. However, this value can be discussed, since the data from Qaim and Döhler (1984), are not in agreement with most recent ones found in the literature (Hermanne et al., 2010) and with ours (see below). Using Qaim et al. data, a specific activity of 25 kCi/g is calculated. The theoretical specific activity is equal to 82 kCi/g, assuming that tin-117 m is the only nucleus in the final product. The difference is due to the presence of the ground state tin-117 g and other stable or long-lived tin isotopes produced during the irradiation.

1.3. Motivations

The ARRONAX cyclotron (Nantes, France) is an accelerator dedicated to the production of radionuclides. It delivers α particles

at 67.4 MeV with an intensity up to 70 μA (Haddad et al., 2008). In this frame, our study focuses on the measurement of the experimental tin-117 m production cross section up to 67.4 MeV from Cd-nat target. Two data sets have been already published in the literature for the Cd-nat(α ,x) reaction, showing some discrepancies between them (Qaim and Döhler, 1984; Hermanne et al., 2010). The advantage of using Cd-nat as target is its availability in metallic form from the Goodfellow (2015) company, with a good purity and homogeneity. Our measurements will allow to get an additional data set to define the behaviour of this reaction, and to determine the other radionuclides produced during the irradiation. From these data, it will be also possible to extract values from the Cd-116(α ,3n)Sn-117 m reaction since only two of the tin stable isotopes that compose the natural cadmium target (see Table 1) can produce Sn-117 m: Cd-114 and Cd-116. Using these data, it will be possible to calculate the production rate and the specific activity of the final product as a function of the α incident energy from 25 MeV to 65 MeV. For the specific activity calculation, TALYS 1.6 code calculations will be used to infer stable tin isotopes production. This will allow to find the best compromise between production yield and specific activity for tin-117 m.

2. Materials and methods

2.1. Experimental set-up

The production cross section data are obtained using the stacked-foils method (Duchemin et al., 2015; Blessing et al., 1995), which consists of the irradiation of a set of thin foils, grouped as patterns. Each pattern contains a target to produce the isotopes of interest. Each target is followed by a monitor foil to have information on the beam intensity thanks to the use of a reference reaction recommended by the International Atomic Energy Agency (IAEA-NDS, 2015). In our experiment, the monitor foil acts also as a catcher to stop the recoil atoms produced in the target foil. A degrader foil is placed after each monitor foil to change the incident beam energy from one target foil to the next one.

Each foil in the stack has been weighed before irradiation using an accurate scale ($\pm 10^{-5}$ g) and scanned to precisely determine its area. The thickness is deduced from these values, assuming that it is homogeneous over the whole surface. In this work, 10 μm thick copper and aluminium monitor foils, 10 μm thick cadmium target foils and 100–500 μm thick aluminium degrader foils were irradiated. The foils, with an isotopic abundance and chemical purity of 99.9%, 99%, 99.7% and 99%, respectively, were purchased from Goodfellow®.

The ARRONAX cyclotron (Haddad et al., 2008) delivers alpha particle beams with an energy uncertainty of ± 0.61 MeV, as

specified by the cyclotron provider using simulations. The beam line is under vacuum and closed using a 75 μm thick kapton foil. The stacks were located about 6.8 cm downstream in air. The energy through each target and monitor foils has been determined in the middle of the thickness of the foil using the SRIM software (Ziegler et al., 2010). Energy loss in the kapton foil and air have been taken into account in our analysis.

All along the stack, depending on the number of foils, the energy uncertainty calculated using the SRIM software (Ziegler et al., 2010) increases up to ± 2 MeV due to the energy straggling (see Table 2). Three stacks were irradiated to cover the energy range from 65 MeV down to 25 MeV (see Table 2), which corresponds to the energy range of interest for the Cd-116(α ,3n) reaction. Since the incident α beam energy delivered by the cyclotron is fixed, in two cases the first foil of the stack was a degrader foil in order to have a different energy through the first cadmium target foil. In addition, the use of several stacks allows also us to minimize the energy uncertainty in our experiment.

With natural cadmium and α particle as projectile, Sn-117 m can be produced from nuclear reactions on Cd-116 or Cd-114. Cd-114 has its maximum contribution between 10 and 25 MeV. This region was not achievable in our experiments without large uncertainties on the projectile energy (higher than 2 MeV). We have then decided to restrict ourselves to 25 MeV, missing most of the contribution of Cd-114.

Irradiations were carried out with a mean beam intensity between 140 and 200 nA particles during one hour. Irradiation conditions are reported also in Table 2.

For all the experiments, the recommended cross sections (Tárkányi et al., 2001) of the Cu-nat(α ,x)Ga-67 (up to 50 MeV) or Al-27(α ,x)Na-22 (from 50 MeV to 70 MeV) reactions were used to get information on the beam intensity.

There is a drawback in using a relative calculation, regarding the possible change in the recommended values when new more accurate measurements are available. The same problem exists for radioactive constants, γ emission branching ratios, etc. Being aware of this problem, we decided to mention the mean energy crossed by each monitor foil, for which our data have been obtained (see Table 2). If the recommended values change (Tárkányi et al., 2001) in the coming years, the cross section data can be modified accordingly.

During irradiation, an instrumented beam stop is used to control the beam current stability. However, it is not used as a faraday cup with precise intensity measurements, since it is not equipped with an electron suppression device.

The activity measurements in each foils were performed using a high purity germanium detector from Canberra (France) with low-background lead and copper shielding. All foils were counted twice. The first measurements started the day after the irradiation (after a minimum of 15 h cooling time) during one hour, for all target and monitor/catcher foils. The second series of measurements were performed one week after EOB, during a minimum of 24 h (one day) and up to 60 h. Gamma spectra were recorded in a suitable geometry calibrated in energy and efficiency with standard Co-57,60 and Eu-152 gamma sources from LEA-CERCA

Table 1
Isotopic composition of the natural cadmium foils from GoodFellow metals.

Cd-106	Cd-108	Cd-110	Cd-111	Cd-112	Cd-113	Cd-114	Cd-116
1.2%	0.9%	12.4%	12.8%	24.0%	12.3%	28.8%	7.6%

Table 2
Irradiation conditions and energies through the cadmium target foils and monitor foils estimated with SRIM (Ziegler et al., 2010).

Beam energy (MeV)	<Intensity> (nA p.)	Energy points in cadmium (MeV)		Energy points in monitor foils (MeV)		
67.40 (61)	196	65.01 (68)	55.13 (93)	64.58 (69)	54.65 (96)	
67.40 (61)	199	48.92 (108)	35.43 (152)	47.41 (111)	33.06 (155)	
67.40 (61)	143	45.86 (116)	39.51 (137)	32.24 (161)	45.06 (118)	31.16 (169)
67.40 (61)	158	42.10 (125)	29.25 (176)	25.26 (193)	41.26 (127)	28.10 (179) 24.02 (199)

(France). The full widths at half maxima were 1.04 keV at 122 keV (Co-57 γ ray) and 1.97 keV at 1332 keV (Co-60 γ ray). The samples were placed at a distance of 19 cm from the detector which is suitable to reduce the dead time and the effect of sum peaks. The dead time during the counting was always kept below 10%.

2.2. Data processing

All the activity values extracted for the radionuclides produced in the targets were derived from the γ spectra and the nuclear decay data (Kinsey et al., 2016; Ekström and Firestone, 2004), given in Table 6 for both tin and cadmium isotopes, and in Table 7 for indium isotopes. The spectra were analysed using the Fitzpeaks spectroscopy software (FitzPeaks Gamma Analysis and Calibration Software version 3.66, 1981).

The recoil nuclei coming from the cadmium targets have been detected and quantified in the monitor foils located after each target. In our experiment, the monitor foils are also used as catcher foils. The activity detected in each monitor foil has been added to that obtained in the preceding target foil.

By knowing the thickness of the foil precisely and the total activity of each isotope produced in the target, their production cross sections are calculated using the activation formula (1) with the appropriate beam current.

$$\sigma(E) = \frac{Act \cdot A}{\chi \cdot \phi \cdot N_a \cdot M \cdot (1 - e^{-\lambda t})} \quad (1)$$

In Eq. (1), the production cross section σ (mb) of a radionuclide at a given energy depends on its measured activity corrected to the time at the end of irradiation Act (Bq), its decay constant λ (s^{-1}), its atomic mass A (g mol $^{-1}$), its areal density M (g.cm $^{-2}$), its chemical and isotopic abundance χ , the Avogadro constant (N_a), the irradiation duration t (s) and the beam current ϕ (p s $^{-1}$).

In our experiment, and because we use thin foils, each target foil receives the same beam current as the monitor foil placed behind it. It is then possible to define a relative Eq. (2) in which the knowledge of the beam current is no longer necessary. In this equation, the prime parameters are associated with Ga-67 or Na-22, used as isotopes of interest for the recommended cross section, while the others relate to the radionuclide produced in the target.

$$\sigma(E) = \sigma'(E) \cdot \frac{\chi' \cdot Act \cdot A' \cdot M' \cdot (1 - e^{-\lambda' t})}{\chi \cdot Act' \cdot A \cdot M \cdot (1 - e^{-\lambda t})} \quad (2)$$

The cross section uncertainty is estimated with a propagation error calculation. Since all the parameters of Eq. (2) are independent, the total error is expressed as a quadratic sum (see Eq. (3)).

$$\frac{\Delta\sigma}{\sigma} = \sqrt{\left(\frac{\Delta\sigma'}{\sigma'}\right)^2 + \left(\frac{\Delta Act}{Act}\right)^2 + \left(\frac{\Delta Act'}{Act'}\right)^2 + \left(\frac{\Delta M}{M}\right)^2 + \left(\frac{\Delta M'}{M'}\right)^2} \quad (3)$$

The main error sources come from the recommended cross section values, the activity value of each produced radionuclide and the areal density uncertainties. The contribution of the irradiation time uncertainty is not significant and has been neglected. Since no uncertainty is given for the recommended cross section values, we have decided to use the uncertainty of the nearest experimental value used by the IAEA to perform the adjustment. It leads to an uncertainty of 11% in average. The uncertainties on the activity of the radionuclides produced in the cadmium targets depend on different parameters such as the gamma line branching ratio, half-life, etc., described afterwards. These uncertainties are, in average, of 1.6% for the tin-117 m activity, 1.9% for the Ga-67 activity and 10.5% for the Na-22. Around 1% of uncertainty is calculated for the areal density.

2.3. Thick target production yield (TTY)

Using the cross section values as a function of the energy $\sigma(E)$ obtained either in this work or in database, we have calculated the associated Thick Target production Yields in MBq for one hour of irradiation and 1 μ Ae, also called 1 h-1 μ A yield. The values are obtained as a function of the projectile energy, using the following expression:

$$TTY = \phi \cdot \chi \cdot \frac{N_a \cdot \rho}{A} (1 - e^{-\lambda \cdot t}) \int_{E_{min}}^{E_{max}} \frac{\sigma(E)}{dE} dE \quad (4)$$

In relation (4), ϕ represents the number of particles per second delivered in one μ Ae. The irradiation time, t , is set at one hour. χ corresponds to the isotopic abundance and chemical purity of the target, ρ is the target density (g.cm $^{-3}$) and N_a is the Avogadro number. $\frac{dE}{dx}$ is the linear energy transfer of the projectile in the target material (MeV cm $^{-1}$). In a thick target, the incident particle energy decreases with the penetration depth. E_{max} corresponds to the incoming projectile energy, whereas E_{min} corresponds to its energy after the target.

2.4. Comparison with the talys 1.6 code

In this work, all the experimental cross section values are compared with the latest version (1.6) of the TALYS code released in December 2013 (Koning and Rochman, 2012). TALYS is a nuclear reaction program which simulates reactions induced by light particles on nuclei heavier than carbon. It incorporates theoretical models to predict observables including cross section values as a function of the incident particle energy (from 1 keV to 1 GeV). A combination of models that best describes the whole set of available data for all projectiles, targets and incident energies have been defined by the authors and put as default in the code. In this way, a calculation can be performed with minimum information in the input file: the type of projectile and its incident energy, the target type and its mass. The results are plotted as TALYS 1.6 default in the following figures. Since there are some differences between experimental data and the results of the TALYS code using default models, we have defined a combination of models, already included in the TALYS code, that better describes the production cross sections, for a variety of projectiles, incident energies and target masses.

The description of the optical, preequilibrium and level density models have been found to have a great influence on the calculated production cross section values. Better results are, in general, obtained when alpha particles are used as projectile using the optical model described by Demetriou et al. (2002), a pre-equilibrium model based on the exciton model including numerical transition rates with optical model for collision probabilities (Gadioli and Hodgson, 1992) and a model for the microscopic level density from Hilaire's combinatorial tables (Goriely et al., 2008). The results of this combination of models are referenced in the following figures as TALYS 1.6 Adj.

3. Results and discussions

This part shows the experimental production cross section results for the Cd-nat(α ,x) reactions. The determination of the Cd-116(α ,3n)Sn-117 m production cross section values and Sn-117 m specific activity is then detailed.

3.1. The $\text{Cd-nat}(\alpha, x)\text{Sn-117 m}$ production cross section

Tin-117 m ($T_{1/2} = 13.60$ (4) d) decays by Isomeric Transition (IT) to its stable ground state tin-117 g by emitting γ rays at 156 and 158 keV (see Table 6). The activity extracted from both γ lines were in good agreement. Our tin-117 m production cross section results are plotted in Fig. 1 as full points.

As reported in Table 6, tin-117 m is produced in the natural cadmium material from the $\text{Cd-114}(\alpha, n)$ reaction with an energy threshold of 5.4 MeV and from the $\text{Cd-116}(\alpha, 3n)$ reaction with an energy threshold of 20.8 MeV. Our results show a maximum around 80 mb and 35 MeV, mainly coming from the $\text{Cd-116}(\alpha, 3n)$ reaction. The peak at low energy, which is related to Cd-114, is not visible in our data set due to our minimal alpha particle energy of 25 MeV.

Two other data sets have been published in the literature (Qaim and Döhler, 1984; Hermanne et al., 2010). The one published in 1984 gives values in the energy range from 15 MeV to 137 MeV and the other one, published in 2010, in the energy range from 5 MeV to 38.5 MeV. These series are shown in Fig. 1 as rhombs and boxes, respectively, up to 80 MeV. Qaim and Döhler (1984), made a high energy range study and Fig. 1 shows only the lower energy part. The points of Qaim and Döhler have been degraded rather far from the initial energy, which explains the energy shift of around 10 MeV in comparison to Hermanne et al.

Our data are in good agreement with those published by Hermanne et al. (2010), and with the energy thresholds of the reactions contributing to the tin-117 m production. Our new set of data complements the trend obtained with the experimental data points from Hermanne et al. (2010). The knowledge of the $\text{Cd-nat}(\alpha, x)\text{Sn-117 m}$ excitation function is then better described between 38 and 65 MeV. With our data and those published in 2010, the contribution from both Cd-114 and Cd-116 to the tin-117 m production with a natural cadmium target are well described.

The experimental data are compared with the results of the TALYS code version 1.6 with default (full line) and adjusted (dash line) models. As an overall result, the TALYS code confirms the trend exhibited by the experimental data. The results from TALYS 1.6 Adj. well follow the experimental trend in energy and amplitude, whereas TALYS 1.6 Default shows an energy shift of a few MeV toward low energies.

3.2. The tin-117 m contaminants production cross sections

3.2.1. The Sn-110 production cross section

Tin-110 has a half-life of 4.154 (4) h (IAEA-NDS, 2015). It decays

by electronic capture to the metastable state of indium-110 ($T_{1/2} = 69.1$ (5) min), by emitting an intense gamma ray of 280.459 keV (97.06 (8)%). This gamma line was used to determine its activity in the recorded gamma spectrum. Tin-110 is produced by (α, xn) reactions on different isotopes in the natural cadmium target, from Cd-108 to Cd-113 in our energy range of interest (see Table 6). Our data are presented in Fig. 2 and Table 8.

Our results are the first ones, for this isotope, above 38 MeV. Below this energy, the agreement with Hermanne et al. (2010), is good. TALYS is able to reproduce the trend. Moreover, the amplitude is well reproduced with TALYS 1.6 Adj.

3.2.2. The Sn-113 production cross section

Tin-113 can be produced either in the ground state, Sn-113 g, or in the isomeric state, Sn-113 m. Sn-113 m has a short half-life of 21.4 (4) min. It decays by electronic capture (8.9%) to In-113 g (stable) and by isomeric transition (91.1%) to Sn-113 g. Sn-113 g, on the other hand, has a long half-life of 115.09 (4) days and decays by electronic capture and β^+ emission to In-113, which emits a high intensity gamma line at 391.690 keV. This gamma ray has been used to determine its activity (see Table 6). Tin-113 m has not been detected in our targets. Its half-life is so short that the 15 h cooling time between the end of bombardment and our measurements has not allowed to quantify its remaining activity during our first measurements. The Sn-113 g activity measured is then cumulative, including that of Sn-113 m. After one to two weeks cooling time, and 24–64 h of counting time, the Sn-113 g cumulative activity has been extracted with an uncertainty of 3.9% in average. Our values are presented in Fig. 3 and Table 8.

Our data allow to highlight the end of the $\text{Cd-111}(\alpha, 2n)$ contribution ($E_s = 15.17$ MeV), that of $\text{Cd-112}(\alpha, 3n)$ from 24.90 MeV and $\text{Cd-113}(\alpha, 4n)$ from 31.66 MeV (see Table 6). However, other measurements are needed to better discriminate the other reaction contributions at higher energies. The data in literature (Qaim and Döhler, 1984; Hermanne et al., 2010) are not in agreement. This difference has been already discussed for the tin-117 m production cross section. We observe some differences in the amplitude between our data and those published by Hermanne et al. (2010). However, looking at the article (Hermanne et al., 2010), our results are comparable since in both cases the Sn-113 g production cross section is cumulative and the 391 keV gamma line is used to extract the Sn-113 g(cum.) activity. The experimental results are too dispersed to conclude on the results of both combinations used in the TALYS code.

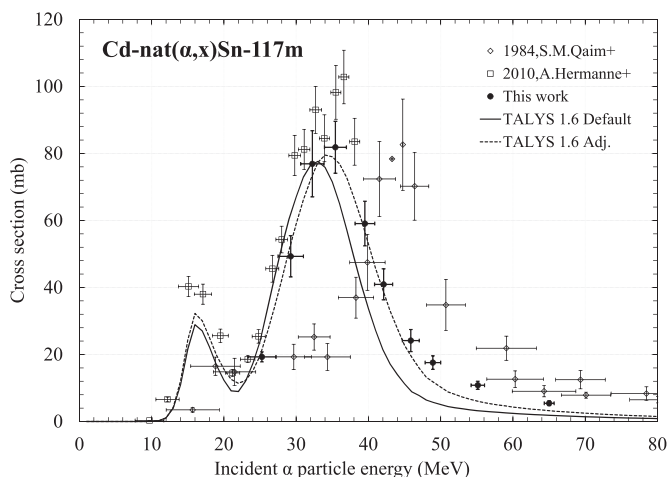


Fig. 1. $\text{Cd-nat}(\alpha, x)\text{Sn-117 m}$ excitation function.

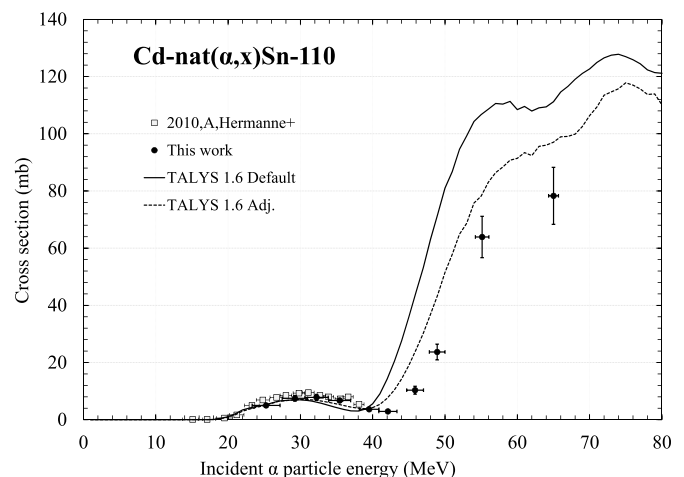


Fig. 2. $\text{Cd-nat}(\alpha, x)\text{Sn-110}$ excitation function.

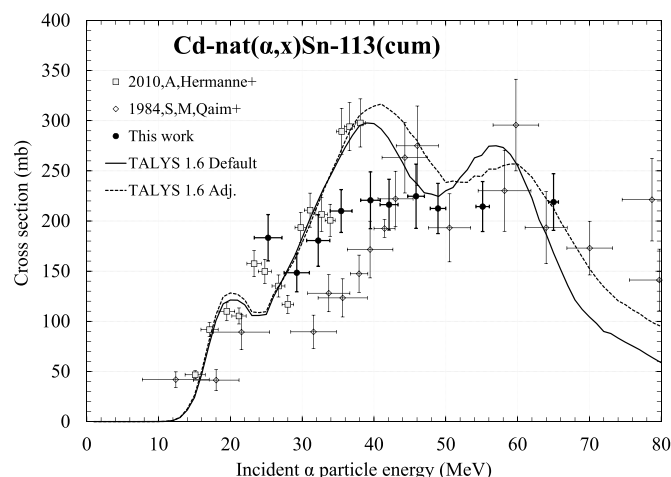


Fig. 3. Cd-nat(α , xn)Sn-113 excitation function (cumulative).

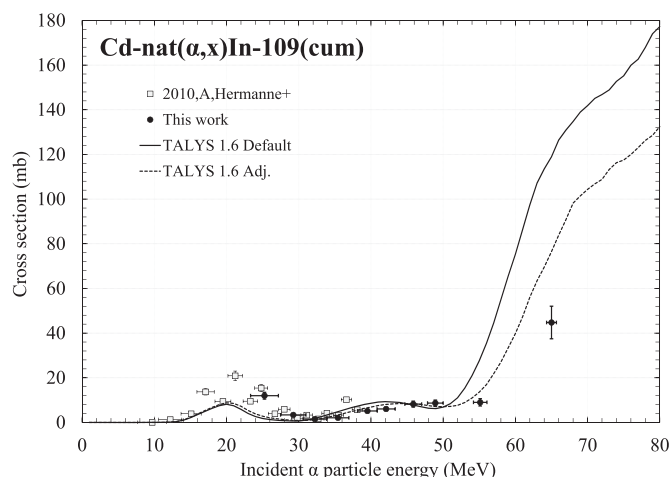


Fig. 4. Cd-nat(α , x)In-109 excitation function (cumulative).

3.2.3. The In-109 production cross section

In-109 is produced by Cd-nat(α , xn+p) reactions. In-109 g ($T_{1/2}=4.2$ (1) h) is also arising from the decay of its metastable states, In-109m2 ($T_{1/2}=0.209$ (6) s) and In-109m1 ($T_{1/2}=1.3$ min), and the decay of Sn-109 ($T_{1/2}=18$ min). The 15 h cooling time between the end of irradiation and the first measurements allowed the decay of the In-109m2, In-109m1 and Sn-109 nuclei produced during the irradiation. The In-109 g activity values obtained from our spectra are then cumulative. In-109 g has been identified and quantified using the 203.5 keV (74%) γ ray emitted from its decay. It emits other γ lines with low branching ratio that led to high uncertainties. These lines were not used in the activity calculation but they allowed to validate the values given by the 203.5 keV γ line. Our data are listed in Table 8 and depicted in Fig. 4.

Our experimental data follow the trend made by the points published in 2010 by Hermanne et al. and give additional information from 38 MeV to 65 MeV. The different reaction contributions (see Table 7) are well defined both by our experimental points and the TALYS code. The TALYS Adj. results are closer to our experimental points than TALYS 1.6 Default, above 55 MeV. Under 30 MeV, TALYS gives underestimated results. Between both energies, the code gives satisfactory results.

3.2.4. The In-110 production cross section

In-110 has a metastable state, In-110 m, with a half-life of 69.1 (5) min. This half-life is too short to obtain information on the produced activity, considering our experimental conditions. In-110 m decays by EC/ β^+ emission to Cd-110 (stable). The activity of the In-110 ground state (In-110 g) has been obtained using the γ lines summarized in Table 7. In-110 g is not subject to the decay of Sn-110, which decays at 100% to In-110 m. Our numerical cross section values are presented in Fig. 5 and Table 9. Our values are in agreement with those published in 2010 by Hermanne et al. Above 50 MeV, additional measurements are needed to better describe the experimental trend. Up to 50 MeV, TALYS 1.6 Adj. gives good results.

3.2.5. The In-111 production cross section

The decay of the indium-111 ground state ($T_{1/2}=2.8047$ (5) days) leads to the emission of two gamma rays (see Table 7) with high branching ratios, which are easily detectable. Cross section results are presented in Fig. 6 and Table 9. They are cumulative values that include contributions from Sn-111 ($T_{1/2}=35.3$ (6) min) and In-111 m ($T_{1/2}=7.7$ (2) min). These results allow to expand our knowledge on the In-111 production with the

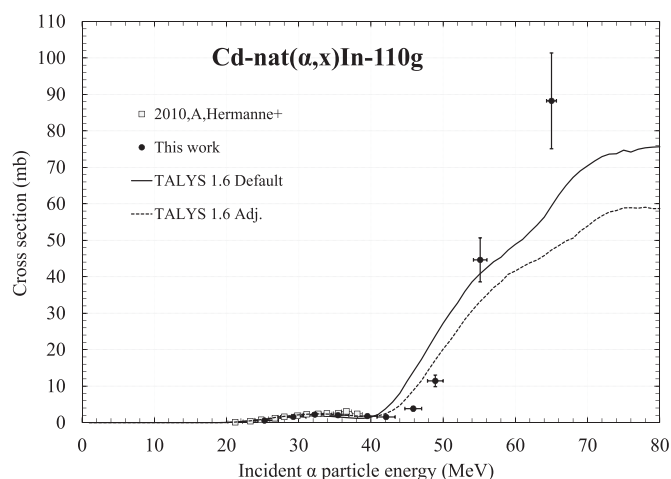


Fig. 5. Cd-nat(α , x)In-110 excitation function.

irradiation of a natural cadmium target by α particles, especially between 40 and 70 MeV. Our results are in agreement with the energy threshold of the different nuclear reactions identified as possible source (see Table 7). There is a small shift in the energy between our data and those of Hermanne et al between 30 MeV and 50 MeV. The TALYS code is able to reproduce the experimental trend.

3.2.6. The In-114 m production cross section

In-114 has two metastable states, In-114m2 ($T_{1/2}=43.1$ (6) s) and In-114m1 ($T_{1/2}=49.51$ (1) d), and an unstable ground state In-114 g ($T_{1/2}=71.9$ (1) s). In-114m2 decays at 100% to In-114m1 by internal transition. The In-114m1 activity has been obtained from the gamma lines summarized in Table 7. In-114 g half-life is 5.6×10^4 times lower than that of In-114m1. Both activities are quickly in secular equilibrium. The cross section values deduced from these activity values are cumulative and are presented in Table 9 and Fig. 7. Our cross section results show a good agreement with those of Hermanne et al. up to 35 MeV. Above this energy, our data are the first to give additional information on the trend. TALYS gives the good trend. Above 55 MeV, TALYS 1.6 Adj. well estimates the excitation function amplitude.

3.2.7. The Cd-115 g production cross section

Cd-115 has a ground state, Cd-115 g ($T_{1/2}=53.46$ (10) h), which is not subject to the decay of its metastable state, Cd-115 m

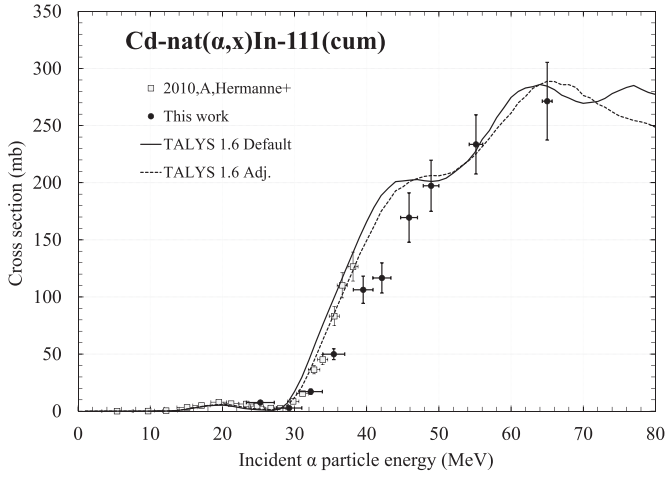


Fig. 6. Cd-nat(α,x)In-111(cum) excitation function.

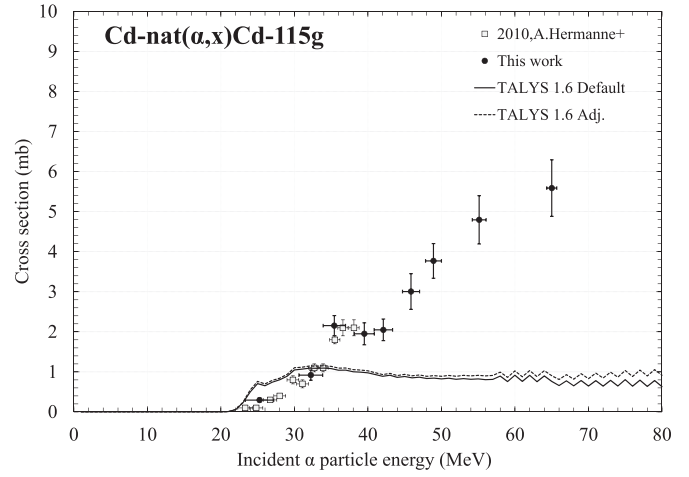


Fig. 8. Cd-nat(α,x)Cd-115g excitation function.

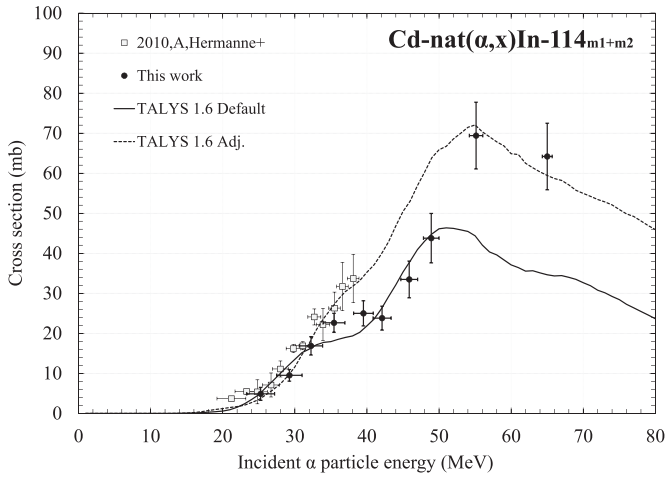


Fig. 7. Cd-nat(α,x)In-114m1+m2 excitation function.

($T_{1/2}=44.6$ (3) days). Cd-115 g decays to In-115 by emitting different gamma rays common with Cd-115 m, even though Cd-115 m has not been quantified due to the low branching ratio of its gamma lines. The γ line at 527.9 keV (see Table 6) was used to determine the Cd-115 g activity since it is the only detectable gamma line different from the Cd-115 m ones. Cross section data are presented in Table 9 and Fig. 8.

Our experimental data well reproduce the ($\alpha, 2p+xn$) reactions contribution, with a shoulder around 40 MeV. Our work is the first one to study the Cd-116($\alpha, 2p+3n$) reaction from threshold (38.27 MeV) up to 65 MeV. Indeed, the values published in 2010 by Hermanne et al. are up to 38 MeV and depends only on the Cd-116($\alpha, \alpha+n$) and Cd-114($\alpha, 2p+n$) reactions. Both sets of data are in good agreement between them. Experimental data are compared with the TALYS code, which is not able to reproduce the experimental trend for these ($\alpha, 2p+xn$) reactions.

3.2.8. The experimental Cd-nat(α,x) reactions production Yields

The production yield values are presented in Fig. 9 for all detected and quantified isotopes in the natural cadmium target. These data have been obtained from Eq. (4) and from our experimental cross section results.

Sn-110, Sn-113 g, In-109, In-110, In-111 are the isotopes which mainly lead to the decrease of the Sn-117 m radionuclidic purity in natural cadmium. At 60 MeV, the In-111 and Sn-110 production yields are, respectively, 17 and 44 times higher than that of Sn-117 m. Indium isotopes can be eliminated by chemical separation.

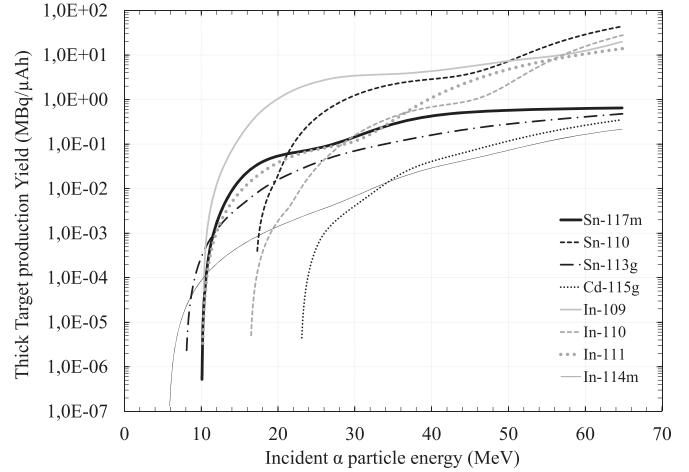


Fig. 9. Production Yield calculated for 1 μ Ae and 1 h of irradiation, from our experimental cross section values obtained for Cd-nat(α,x) reactions.

Sn-110 and Sn-113 g can be avoided using enriched cadmium as target material and appropriate incident beam energy.

3.3. The tin-117 m production from the Cd-116($\alpha, 3n$) reaction

As described in the material and method part, in natural cadmium only Cd-114 and Cd-116 can contribute to the production of tin-117. Cd-114 has its main contribution at low energy as it can be seen from data published by Adam Rebeles et al. (2008) on the energy range from 12 to 38 MeV. The contribution of Cd-114 can be estimated and subtracted from the values obtained from Cd-natural, to get a new evaluation of the Cd-116($\alpha, 3n$) contribution. The data obtained for the Cd-116($\alpha, 3n$) reaction have then been compared with the existing ones, both for cross section and thick target production yield values. Using the TALYS code, all the other tin isotopes produced during the irradiation have been considered including stable tin isotopes, which allows to calculate the specific activity as a function of the projectile energy.

3.3.1. Determination of the Cd-116($\alpha, 3n$)Sn-117 m excitation function

Data exist in the literature for the Cd-114(α, n) excitation function. They have been published by Adam Rebeles et al. (2008) on the energy range from 12 to 38 MeV. Up to the energy threshold of the Cd-116($\alpha, 3n$) reaction, i.e. 20.8 MeV, it is the only contribution for the production of Sn-117 m. The first step was,

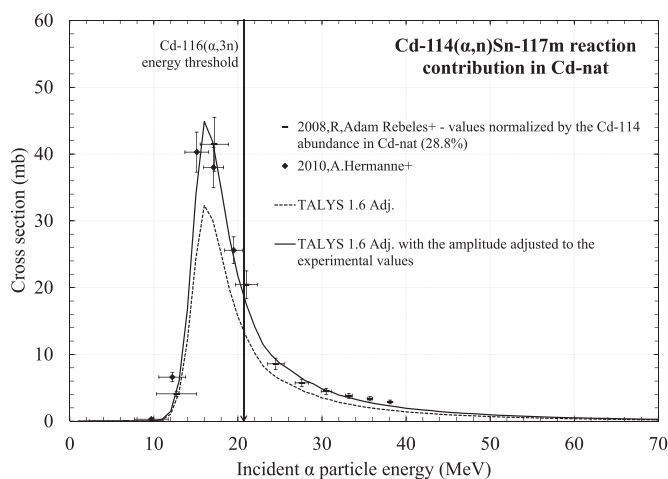


Fig. 10. Cd-114(α ,n)Sn-117 excitation function normalized to a Cd-nat target.

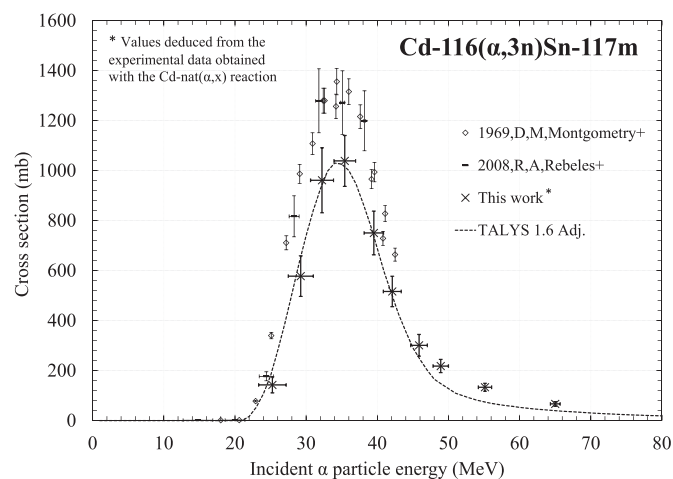


Fig. 11. Cd-116(α ,3n)Sn-117 m excitation function.

therefore, to scale down the data from Adam Rebeles et al. (2008), by multiplying the values by the isotopic abundance of Cd-114 in natural Cd (28.8% – see Table 1). These scaled data are presented in Fig. 10. Data from Hermanne et al. (2010), are also used, up to the Cd-116(α ,3n) energy threshold.

In Fig. 10, values from the TALYS code 1.6 Adj. are presented as dashed line. These values correspond to a natural abundance in Cd-nat of 28.8%. TALYS values are lower than the experimental ones even if they show a similar behaviour. In order to get information on the Cd-114(α ,n) production cross section at higher energy, we have decided to use the TALYS 1.6 adj. results, applying a coefficient to bring these values close to the experimental data.

In Fig. 10, the full line shows the behaviour of the normalized values of the TALYS code. The agreement between the data and the normalized value is very good. These data can then be used to calculate the Cd-114(α ,3n)Sn-117 m cross section values up to 65 MeV. These values, called “TALYS 1.6 Adj. – with the amplitude adjusted to the experimental values” in Fig. 10, are used to determine the contribution of the Cd-114(α ,n) reaction from our experimental data obtained using natural cadmium. 13% of uncertainty is considered on the normalized and adapted values from TALYS, corresponding to the larger uncertainty on the experimental data of Adam Rebeles et al. (2008) and Hermanne et al. (2010) used to fix the TALYS trend. After subtraction of this contribution from our Cd-nat(α ,xn)Sn-117 m cross section results, the obtained data are normalized to take into account the fact that Cd-116 represents only 7.6% of the atoms in the natural cadmium target. Our calculated data are plotted as crosses in Fig. 11 and the numerical values are listed in Table 3.

Our values for the Cd-116(α ,3n)Sn-117 m cross section show a maximum at 36 MeV of around 1 barn. The two published set of data (Montgomery and Porile, 1969; Adam Rebeles et al., 2008) show similar results with a maximum around 36 MeV. However, in comparison with our data, their maximum value is 13% higher than ours. The authors of the experiments published in 1969 and 2008 have determined the Cd-116(α ,3n)Sn-117 m production cross section using electroplating method for the target preparation, and enriched Cd-116 material (respectively 97.2% and 97.7%).

For the Cd-nat(α ,x) reaction, the cross section values from our work and those of Hermanne et al. (2010), show the same amplitude. Data from Adam Rebeles et al. (2008), normalized for a Cd-nat target are 20% higher. Furthermore, in the article published by Montgomery and Porile (1969), the γ lines used to extract the Sn-117 m activity are presented at 159 and 161 keV. Today, the energy values tabulated in the nuclear data bases (Kinsey et al., 2016; Ekström and Firestone, 2004) are different. Their data will

Table 3

Tin-117 m production cross section values from the Cd-116(α ,3n), deduced from our experimental data obtained for the Cd-nat(α ,x)Sn-117 m reaction.

Energy (MeV)	σ Sn-117 m (mb)
65.01 \pm 0.68	66.61 \pm 9.18
55.13 \pm 0.93	133.23 \pm 15.95
48.92 \pm 1.08	218.20 \pm 26.59
45.86 \pm 1.16	301.13 \pm 43.41
42.10 \pm 1.25	516.54 \pm 61.05
39.51 \pm 1.37	750.52 \pm 87.69
35.43 \pm 1.52	1038.63 \pm 101.98
32.24 \pm 1.61	961.22 \pm 130.13
29.25 \pm 1.76	577.68 \pm 81.51
25.26 \pm 1.93	142.56 \pm 35.05

also strongly depend on the target homogeneity and the Cd-116 isotopic purity.

In our case, our calculation strongly depends on the experimental data obtained with the Cd-nat target and on the adjusted TALYS values for the Cd-114(α ,n) reaction (see Fig. 10). The values obtained in the energy range between 25 and 30 MeV are the most dependent to the Cd-114(α ,n) reaction, as shown in Fig. 10.

Our Cd-116(α ,3n)Sn-117 m excitation function results shown in Fig. 11 are in good agreement with the TALYS 1.6 Adj. results. They give the same energy and value for the maximum of the cross section.

3.3.2. The tin-117 m thick Cd-116 target production yield

The Thick Target production Yield (TTY) is calculated using the relation shown in part 2.3 after applying a spline interpolation on the experimental data points. First, a fit from the Montgomery and Porile (1969) and Adam Rebeles et al. (2008), cross section values has been done. From this fit, the thick target production yield has been calculated and is presented in Fig. 12 as a function of the incident α particles energy. It reaches 6.1 MBq/ μ A h at 42.5 MeV, the maximal energy available with experimental data. Secondly, a fit is applied to our data assuming a 100% Cd-116 enriched target. The thick target production yield reaches 4.4 MBq/ μ A h at 42.5 MeV (see Fig. 12), which corresponds to 72% of the value obtained with the literature data (Montgomery and Porile, 1969; Adam Rebeles et al., 2008).

The Clear Vascular company produces tin-117 m using the Cd-116(α ,3n) reaction with 99% Cd-116 enriched target. The corresponding Thick Target production Yield over the energy range

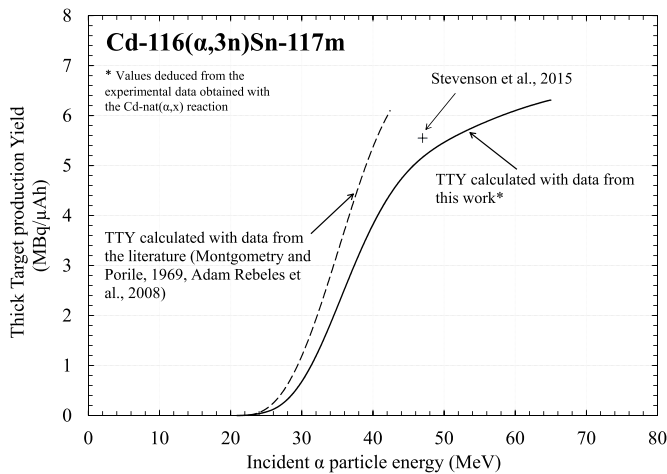


Fig. 12. Cd-116($\alpha,3n$)Sn-117 m production Yield calculated for 1 μ Ae and 1 h of irradiation.

47 MeV down to 20 MeV is consistent with 150 μ Ci/ μ A h, or 5.55 MBq/ μ A h (Stevenson et al., 2015), determined using the data published in Qaim and Döhler (1984). In one hand, for the same energy range, our work gives a value of 5.2 MBq/ μ A h considering a 100% Cd-116 enriched target. This value is close to that presented by the Clear Vascular Company (Stevenson et al., 2015). In the other hand, the yield obtained from the data published in 1969 and 2008 (Montgomery and Porile, 1969; Adam Rebeles et al., 2008) overestimates the value of Stevenson et al. (2015), for a reduced energy range from 42.5 MeV down to 20 MeV. Using a 65 MeV α particle beam with a Cd-116 target thick enough to decrease the incident energy down to 20.80 MeV (the Cd-116($\alpha,3n$) energy threshold), the production yield reaches 6.3 MBq/ μ Ah from our data (see Fig. 12).

3.4. The tin-117 m radionuclidic purity

Considering the production of other radioactive tin isotopes during the irradiation of a 100% Cd-116 target, Sn-110 and Sn-111 will not be produced, since the energy threshold is higher than the energy considered in our study (65 MeV). However, tin-113 can be produced with the Cd-116($\alpha,7n$) reaction from its energy threshold at 56.3 MeV. Tin-113 can be totally avoided with α particle energy beam under 56 MeV. With 56 MeV, the Sn-117 m production yield reaches 92% of that with 65 MeV, which corresponds to 5.8 MBq/ μ Ah. The best radioisotopic purity will be obtained, in this case, with 56 MeV α beam.

3.5. The tin-117 m specific activity (SA)

3.5.1. The specific activity (SA)

The Specific Activity (SA) is the ratio between the activity of the radionuclide of interest and the total mass of all isotopes of the same element standing in the sample. The specific activity is usually expressed in term of activity per element quantity: Ci/mol, Ci/g, Bq/mol, Bq/g or MBq/nmol (see Eq. (5)).

$$SA = \frac{Act_{\text{isotope of interest}}}{m_{\text{isotopes of the same element}}} \quad (5)$$

If all atoms standing in the sample are those of the radionuclide of interest, the SA is maximum. However, the irradiation of a target usually leads to the production of other stable or radioactive isotopes of the same element. In these cases, the SA value decreases. The SA is of relevance for radiolabeling and for radio-pharmaceuticals, for the determination of their chemical and/or

biological effect on the targeted system (IAEA, 2008). Indeed, if there are other isotopes than the one of medical interest, some vector molecules are labelled with these other isotopes, which will be inefficient.

With cyclotrons it is possible to reach high specific activities since the use of charged particles allows to produce a radionuclide of interest for which its chemical element is different to the one composing the target.

3.5.2. The Sn-117 m SA determination method and discussion

For the tin-117 m SA determination, the Sn-117 m production yield is needed. This value has been determined previously from our experimental data. In addition, the mass of all tin isotopes is needed. First, we suppose that a chemical separation is made quickly after the EOB and that all the remained isotopes in the product are Sn isotopes. Using a 65 MeV beam, Sn-113 to Sn-120 are produced. No data are available for stable isotopes. The production cross section values of all tin isotopes that can be produced with a 65 MeV α beam on a Cd-116 target have been determined using TALYS 1.6 Adj.

The SA is calculated at the EOB up to 65 MeV using the number of Sn-114,115,116,117 g,118,119 g,120 stable atoms and that of Sn-113 m ($T_{1/2}=21.4$ (4) min), Sn-113 g ($T_{1/2}=115.09$ (4) days) and Sn-119 m ($T_{1/2}=293.1$ (7) days). The values are plotted in Fig. 13 in kCi/g, as a function of the incident α particles energy. At EOB, the tin-117 m SA reached 41.4 kCi/g at 39.4 MeV.

The mass contribution (%) of each Sn isotope is shown in

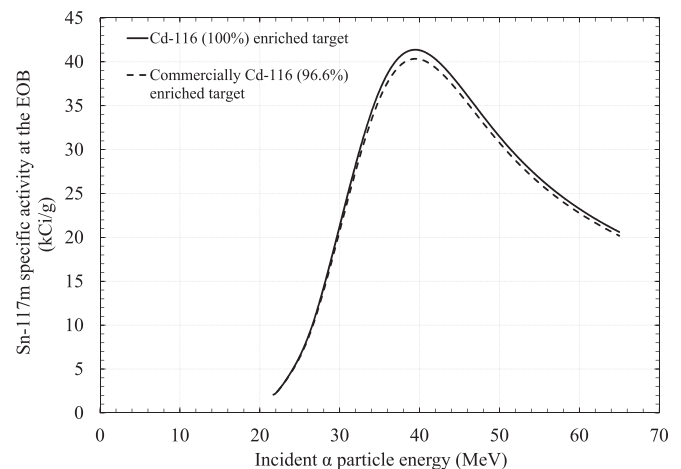


Fig. 13. Tin-117 m specific activity at the EOB calculated from a 100% Cd-116 target (full line) and a commercialized Cd-116 enriched target (dash line), as a function of the incident α particles energy.

Table 4

Mass contribution in % of the Sn isotopes in the sample at EOB, for a Cd-116 target irradiated by different beam energies down to the Cd-116($\alpha,3n$) reaction threshold.

Energy	30 MeV	40 MeV	50 MeV	60 MeV
Sn-113 m	0	0	0	0.0012
Sn-113 g	0	0	0	0.00021
Sn-114	0	0	0.0050	1.71
Sn-115	0	0.0036	4.85	20.91
Sn-116	0.017	11.71	34.67	33.70
Sn-117 g	6.43	5.42	3.47	2.54
Sn-117 m	25.95	50.38	38.40	28.36
Sn-118	66.31	31.75	18.12	12.44
Sn-119 m	0.58	0.33	0.22	0.16
Sn-119 g	0.72	0.41	0.26	0.19
Sn-120	0.000092	0.000034	0.000018	0.000012

Table 4, for different energies impinging a Cd-116 target thick enough to decrease the initial energy down to the Cd-116($\alpha,3n$) energy threshold.

3.5.3. The Sn-117 m SA from a commercialized Cd-116 enriched product

Trace Sciences International Inc. (2015), provides enriched material including Cd-116, with an isotopic composition presented in Table 5.

Such information allows to calculate the specific activity based on a commercialized Cd-116 material, still starting from the Cd-116 ($\alpha,3n$) energy threshold.

Table 5
Isotopic composition of the Cd-116 enriched product provided by Trace Sciences.

Cd-106	Cd-108	Cd-110	Cd-111	Cd-112	Cd-113	Cd-114	Cd-116
0.01%	0.01%	0.25%	0.25%	0.71%	0.38%	1.79%	96.6%

Based on the fact that the amount of atoms of Cd-106, 108, 110 and Cd-111 is negligible (less than 0.3%), only nuclear reactions on Cd-112,113,114 and Cd-116 are considered. Cd-114 and Cd-116 are the only Cd isotopes leading to the production of Sn-117 m. The Sn-117 m production yield obtained from our experimental data for a 100% Cd-116 enriched target (see Fig. 12) is then normalized by the real percentage of Cd-116; 96.6% (see Table 5). For the Cd-114(α,n) reaction the Sn-117 m production is calculated from the production cross section of TALYS 1.6 Adj. normalized as presented in Fig. 10, and reduced by the percentage of Cd-114 in the enriched product. Both values are added.

The other Cd isotopes composing the enriched material lead to the decrease of the specific activity and radionuclidic purity by producing other Sn isotopes atoms. Based on the same method, previously presented, the number of atoms produced from the different Cd isotopes composing the enriched material has been calculated using the TALYS 1.6 Adj. code, normalized by their respective percentage.

This calculation gives a maximum Sn-117 m specific activity at 39.5 MeV and 40.3 kCi/g, close to that obtained assuming a 100%

Table 6
Physical characteristics of tin and cadmium radionuclides (QCALC, 1995; Kinsey et al., 2016; Ekström and Firestone, 2004).

Radionuclide	$T_{1/2}$	E_γ (keV)	I_γ (%)	Contributing reaction (s)	Threshold (MeV)
Sn-117 m	13.60 (4) d	156.02	2.113 (6)	Cd-114(α,n)	5.45
		158.562	86	Cd-116($\alpha,3n$)	20.80
Sn-110	4.154 (4) h	280.459	97.06 (8)	Cd-108($\alpha,2n$)	17.76
				Cd-110($\alpha,4n$)	35.62
				Cd-111($\alpha,5n$)	42.83
				Cd-112($\alpha,6n$)	54.52
				Cd-113($\alpha,7n$)	63.47
Sn-113 g	115.09 (4) d	391.690	64	Cd-110(α,n)	7.95
				Cd-111($\alpha,2n$)	15.17
				Cd-112($\alpha,3n$)	24.90
				Cd-113($\alpha,4n$)	31.66
				Cd-114($\alpha,5n$)	41.01
				Cd-116($\alpha,7n$)	56.34
Cd-115 g	53.46 (10) h	527.9	27.45 (18)	Cd-116($\alpha,\alpha+n$)	9.00
				Cd-114($\alpha,2p+n$)	22.93
				Cd-116($\alpha,2p+3n$)	38.27

Table 7
Physical characteristics of indium radionuclides (QCALC, 1995; Kinsey et al., 2016; Ekström and Firestone, 2004).

Radionuclide	T _{1/2}	E _γ (keV)	I _γ (%)	Contributing reaction (s)	Threshold (MeV)
In-109	4.2 (1) h	203.5	74	Cd-106(α,p)	5.72
				Cd-110(α,2n + p)	24.65
				Cd-110(α,4n + p)	42.50
				Cd-111(α,5n + p)	49.72
				Cd-112(α,6n + p)	59.43
In-110	4.9 (1) h	641.68	25.9 (6)	Cd-108(α,n + p)	16.30
		707.4	29.5 (10)	Cd-110(α,3n + p)	34.16
		937.493	68.4 (14)	Cd-111(α,4n + p)	41.37
		997.256	10.52 (20)	Cd-112(α,5n + p)	51.09
				Cd-113(α,6n + p)	57.84
In-111	2.8047 (5) d	171.28	90	Cd-108(α,p)	5.94
		245.395	94	Cd-110(α,2n + p)	23.80
				Cd-111(α,3n + p)	31.02
				Cd-112(α,4n + p)	40.74
				Cd-113(α,5n + p)	47.50
				Cd-114(α,6n + p)	56.85
In-114m1	49.51 (1) d	190.29	15.56 (15)	Cd-111(α,p)	6.25
		558.456	3.24 (23)	Cd-112(α,n + p)	15.98
		725.298	3.24 (23)	Cd-113(α,2n + p)	22.74
				Cd-114(α,3n + p)	32.10
				Cd-116(α,5n + p)	47.43

Table 8Production cross section values of Sn-110, Sn-113 and Sn-117 m from the Cd-nat(α , xn) reaction and In-109 from the Cd-nat(α , x) reaction.

Energy (MeV)	σ Sn-117 m (mb)	σ Sn-113 (cum.) (mb)	σ Sn-110 (mb)	σ In-109 (mb)
65.01 \pm 0.68	5.44 \pm 0.70	218.91 \pm 28.25	78.29 \pm 9.97	44.74 \pm 7.31
55.13 \pm 0.93	10.82 \pm 1.21	214.40 \pm 24.94	63.90 \pm 7.23	8.99 \pm 1.63
48.92 \pm 1.08	17.58 \pm 2.02	212.50 \pm 25.01	23.68 \pm 2.75	8.67 \pm 1.37
45.86 \pm 1.16	24.16 \pm 3.29	224.67 \pm 32.06	10.32 \pm 1.38	8.21 \pm 1.34
42.10 \pm 1.25	40.93 \pm 4.63	216.31 \pm 25.30	2.89 \pm 0.34	6.03 \pm 0.71
39.51 \pm 1.37	59.07 \pm 6.66	220.67 \pm 28.33	3.63 \pm 0.44	5.14 \pm 0.73
35.43 \pm 1.52	81.84 \pm 7.74	209.97 \pm 21.38	6.71 \pm 0.70	2.07 \pm 0.34
32.24 \pm 1.61	76.89 \pm 9.88	180.53 \pm 25.70	7.80 \pm 1.03	1.51 \pm 0.26
29.25 \pm 1.76	49.29 \pm 6.15	148.41 \pm 19.00	7.48 \pm 0.98	3.38 \pm 0.52
25.26 \pm 1.93	19.28 \pm 2.43	183.31 \pm 23.10	4.99 \pm 0.64	11.97 \pm 1.54

Table 9Production cross section values of In-110 g, In-111, In-114 m and Cd-115 g from the Cd-nat(α , x) reaction.

Energy (MeV)	σ In-110 g (mb)	σ In-111 (mb)	σ In-114 m (mb)	σ Cd-115 g (mb)
65.01 \pm 0.68	88.23 \pm 13.14	271.38 \pm 34.00	64.18 \pm 8.32	5.59 \pm 0.71
55.13 \pm 0.93	44.62 \pm 6.04	233.57 \pm 25.90	69.40 \pm 8.34	4.80 \pm 0.60
48.92 \pm 1.08	11.44 \pm 1.59	197.33 \pm 22.34	43.78 \pm 6.17	3.77 \pm 0.43
45.86 \pm 1.16	3.83 \pm 0.61	169.51 \pm 21.64	33.47 \pm 4.59	3.00 \pm 0.45
42.10 \pm 1.25	1.53 \pm 0.61	116.65 \pm 13.19	23.80 \pm 2.99	2.05 \pm 0.27
39.51 \pm 1.37	1.78 \pm 0.27	106.31 \pm 11.94	24.99 \pm 3.15	1.95 \pm 0.27
35.43 \pm 1.52	2.03 \pm 0.29	49.91 \pm 4.73	22.63 \pm 2.37	2.15 \pm 0.25
32.24 \pm 1.61	2.20 \pm 0.35	17.11 \pm 2.20	16.87 \pm 2.28	0.92 \pm 0.13
29.25 \pm 1.76	1.56 \pm 0.26	2.77 \pm 0.35	9.51 \pm 1.46	
25.26 \pm 1.93	0.55 \pm 0.08	7.57 \pm 0.95	4.89 \pm 1.62	0.30 \pm 0.05

Cd-116 target (i.e. 39.4 MeV and 41.4 kCi/g). With the same target, the Sn-117 m production yield reaches 6.1 MBq/ μ A h at 65 MeV, against 6.3 MBq/ μ A h with a 100% Cd-116 target.

However, this isotopic composition reduces the radionuclidic purity by the production of the long half-life Sn-113 g, that can be produced from 24.50 MeV with the Cd-112(α ,3n) reaction. Sn-113 g is also produced below this energy if we consider the reactions on Cd-106 to Cd-111 atoms. Sn-113 has a metastable state, Sn-113 m, that decays by isomeric transition to Sn-113 g. Few hours after the EOB, Sn-113 m has totally decayed and 91.1% of the Sn-113 m atoms have been transformed to Sn-113 g atoms (see part 3.2.2). At that time and considering only the reactions on Cd-112,113,114 and Cd-116, the Sn-113 g activity represents 0.16% of the Sn-117 m activity, against 0.008% at the EOB (without Sn-113 m decay).

4. Conclusion

Tin-117 m is a radionuclide of medical interest, already used in pre-clinical and clinical trials for vulnerable plaque. Previous studies have shown that the Cd-116(α ,3n) reaction leads to the highest purity product, with a production rate in agreement with that required for medical applications. Due to the lack of data at an energy higher than 42.5 MeV, tin-117 m and contaminants cross section measurements have been investigated at the ARRONAX cyclotron using natural cadmium targets and alpha particles up to 65 MeV. From these results, a method has been defined to extract the tin-117 m production cross section values from a Cd-116 enriched target. Thick target yield, radionuclidic purity and specific activity have been studied using experimental data and the TALYS 1.6 code values. This latter step was necessary for the determination of the number of stable and long-lived isotopes produced during the irradiation. Using the isotopic composition of commercialized enriched Cd-116 target, the maximum specific activity

reached 40.3 kCi/g at 39.5 MeV. From our work, the irradiation parameters can be determined as a function of the required purity of the final product from 20.8 MeV up to 65 MeV.

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