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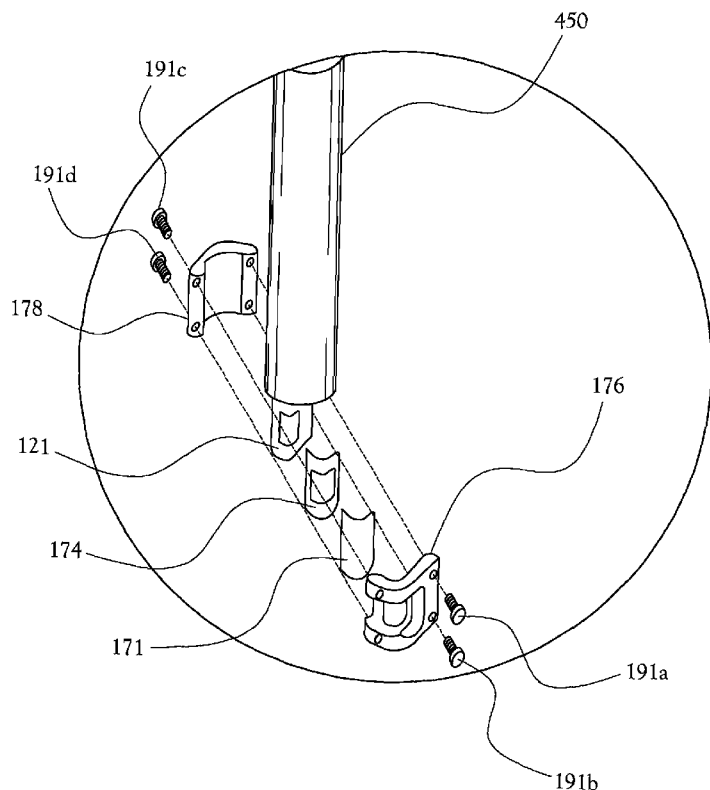
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(54) Title: RADIOISOTOPE TARGET ASSEMBLY



(57) Abstract: A target assembly to produce radioisotopes for the synthesis of radiopharmaceuticals. The target assembly includes a target vessel with a target chamber adapted to receive a target material. A thin cover sheet of particle-permeable material covers the target chamber. In a bombardment process, a high-energy particle beam generated by a cyclotron or particle accelerator strikes the thin cover sheet, whereby at least some of the particles from the particle beam penetrate to the target chamber so as to interact with the target material, altering the nuclear makeup of some of the atoms in the target material to produce radioisotopes.



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TITLE OF INVENTION

Radioisotope Target Assembly

CROSS-REFERENCE TO RELATED APPLICATIONS

[0001] This application claims the benefit of United States Patent Application No. 13/248,906, filed on September 29, 2011, which is incorporated herein in its entirety by reference.

STATEMENT REGARDING FEDERALLY-SPONSORED
RESEARCH OR DEVELOPMENT

Not Applicable

BACKGROUND OF THE INVENTION

1. Field of Invention

[0002] The present general inventive concept relates to an apparatus and method to bombard a nucleus with charged particles so as to bring about a change in the nucleus resulting in a different isotope of the original nucleus or in a different element; and more particularly, to an apparatus to position a target material in the path of a stream of charged particles in order to produce a radioisotope for use in a radiopharmaceutical.

2. Description of the Related Art

[0003] A biomarker is used to interrogate a biological system and can be created by “tagging” or labeling certain molecules, including biomolecules, with a radioisotope. A biomarker that includes a positron-emitting radioisotope is required for positron-emission tomography (PET), a noninvasive diagnostic imaging procedure that is used to assess perfusion or metabolic, biochemical and functional activity in various organ systems of the human body. Because PET is a very sensitive biochemical

imaging technology and the early precursors of disease are primarily biochemical in nature, PET can detect many diseases before anatomical changes take place and often before medical symptoms become apparent. PET is similar to other nuclear medicine technologies in which a radiopharmaceutical is injected into a patient to assess metabolic activity in one or more regions of the body. However, PET provides information not available from traditional imaging technologies, such as magnetic resonance imaging (MRI), computed tomography (CT) and ultrasonography, which image the patient's anatomy rather than physiological images. Physiological activity provides a much earlier detection measure for certain forms of disease, cancer in particular, than do anatomical changes over time.

[0004] A positron-emitting radioisotope undergoes radioactive decay, whereby its nucleus emits positrons. In human tissue, a positron inevitably travels less than a few millimeters before interacting with an electron, converting the total mass of the positron and the electron into two photons of energy. The photons are displaced at approximately 180 degrees from each other, and can be detected simultaneously as "coincident" photons on opposite sides of the human body. The modern PET scanner detects one or both photons, and computer reconstruction of acquired data permits a visual depiction of the distribution of the isotope, and therefore the tagged molecule, within the organ being imaged.

[0005] Most clinically-important positron-emitting radioisotopes are produced in a cyclotron. Cyclotrons operate by accelerating electrically-charged particles along outward, quasi-spherical orbits to a predetermined extraction energy generally on the order of millions of electron volts. The high-energy electrically-charged particles form a continuous beam that travels along a predetermined path and bombards a target. When the bombarding particles interact in the target, a nuclear reaction occurs at a sub-atomic level, resulting in the production of a radioisotope. The radioisotope is then combined chemically with other materials to synthesize a radiochemical or radiopharmaceutical suitable for introduction into a human body.

[0006] Figures 1 and 2 depict a conventional cyclotron used for the production of radioisotopes. As shown in Figure 2, the cyclotron **6** includes an array of four “D” electrodes, also known as “dees” **61**. The dees **61** are positioned in the valleys **62** of a large electromagnet **63**. As shown in Figure 1 and in the exploded view of the same cyclotron in Figure 2, during operation of the cyclotron **6**, an ion source continuously generates charged particles and introduces them into the cyclotron **6** at the center of the array of dees **61**. The charged particles are exposed to a strong magnetic field generated by opposing magnet poles situated above and below the array of dees **61**. A radio frequency (RF) oscillator applies a high frequency, high voltage signal to each of the dees **61** causing the charge of the electric potential developed across each of the dees **61** to alternate at a high frequency. Neighboring dees **61** are given opposite charges such that charged particles entering the gap between neighboring dees **61** see a like charge on one neighboring dee **61** and an opposite charge on the other neighboring dee **61**, which results in acceleration (i.e., increasing the energy) of the charged particles. With each energy gain, the orbital radius of the charged particles increases. The result is a stream of charged particles **A** following an outwardly spiraling path away from the center of the array of dees **61**. The charged particles ultimately exit the cyclotron **6** as a particle beam **B** directed at a target **11**.

[0007] As shown in Figures 1 and 2, the particle beam **B** leaves the magnetic field of the cyclotron **6** before passing through a beam tube **91** and a collimator **93** to strike a target **11**. The beam tube **91** and collimator **93** help to keep the particle beam focused after it leaves the magnetic field of the cyclotron **6**. Figure 3 shows an exploded view of a cyclotron **6'** in use with an internal target **11'**—that is, a target that is positioned within the magnetic field of the electromagnet **63**, so that a particle beam **B'** generated by the cyclotron **6'** does not need to leave the magnetic field of the electromagnet **63** before striking the target **11'**. Such an internal target has certain advantages over an external target. When using an external target, the particle beam loses some energy and concentrated power as it travels the distance between the cyclotron and the external target. Using an internal target, on the other hand, avoids

this loss of beam energy and focus since the particle beam does not leave the immediate area of the cyclotron. This means that the particle stream generated by the cyclotron need not be as highly energetic as would be the case if it were necessary to compensate for a loss of beam energy and focus over distance. Therefore, using an internal target to position a target material in the path of the particle beam allows for the use of a smaller, less powerful cyclotron, with less attendant radiation and less need for shielding or extensive physical plant. Further, the elimination of the beam tube and collimator results in fewer total components contaminated by radiation.

BRIEF SUMMARY OF THE INVENTION

[0008] The present general inventive concept is directed toward a target assembly for use with a cyclotron in producing radioisotopes for the synthesis of radiopharmaceuticals. The cyclotron accelerates small charged particles, such as protons, deuterons or helium nuclei, to form a high-energy particle beam. The particle beam then strikes a designated target area on the target assembly so as to interact with a target substance (i.e. the “target material”). The interaction with the charged particle beam alters the nuclear makeup of some of the atoms in the target material, thereby producing radioisotopes. These radioisotopes will in short time decay, emitting positrons or other energy signatures in the process. When incorporated into radiopharmaceutical molecules, these radioisotopes have useful medical applications, for instance in positron emission tomography (PET).

[0009] The target assembly includes a target vessel defining a target chamber adapted to receive target material. A thin sheet of particle-permeable material covers the target chamber and is welded to the target vessel. A target material input is provided in fluid communication with the target chamber to deliver a target material to the target chamber. A cooling system is provided in communication with the target assembly. During a bombardment process, the cooling system keeps the

target vessel from overheating while a particle beam from a cyclotron strikes the target material within the target chamber, thereby transforming the target material to contain a radioisotope. A gas supply is provided to keep the target material under pressure in the target chamber during the bombardment process. Following the bombardment process, the transformed target material is evacuated from the target chamber and directed to a chemical processing unit, where at least a portion of the radioisotopes formed within the transformed target material are combined with other reagents to synthesize a radiopharmaceutical.

[0010] In many embodiments of the present invention, the target material used is heavy water —i.e. H_2O molecules in which the oxygen atom consists of the O-18 isotope. Likewise, in many embodiments of the present invention, the radioisotope produced by the bombardment process is the F-18 isotope of fluorine. However, the present invention contemplates the use of other target materials with the present invention, and the production of other radioisotopes.

BRIEF DESCRIPTION OF THE SEVERAL VIEWS OF THE DRAWINGS

[0011] The above-mentioned features of the invention will become more clearly understood from the following detailed description of the invention read together with the drawings in which:

Figure 1 is a schematic illustration of a cyclotron directing a beam of charged particles toward an external target;

Figure 2 is an exploded view of the cyclotron system shown in Figure 1;

Figure 3 is an exploded view of a cyclotron system directing a beam of charged particles toward an internal target;

Figure 4 is a schematic diagram of one embodiment of the target assembly;

Figure 5 is a partial view showing the target vessel component of the target assembly shown in Figure 4;

Figure 6 is a perspective view of another embodiment of the target assembly;

Figure 7 is a close-up look at an exploded view of the target vessel of the same embodiment shown in Figure 6;

Figure 8A is a side view of the embodiment shown in Figure 6;

Figure 8B is another side view of the embodiment shown in Figure 6, showing the target assembly viewed from a different perspective;

Figure 9 is a sectional view of the embodiment shown in Figure 6, taken along the line 9-9 of Figure 8B;

Figure 10 is a sectional view of the embodiment shown in Figure 6, taken along the line 10-10 of Figure 8B; and

Figure 11 is a sectional view of the embodiment shown in Figure 6, taken along the line 11-11 of Figure 9.

DETAILED DESCRIPTION OF THE INVENTION

[0012] A target assembly for use with a cyclotron or accelerator in producing radioisotopes for the synthesis of radiopharmaceuticals is described more fully herein. The invention may be embodied in many different forms and should not be construed as limited to the embodiments set forth herein.

[0013] Figure 4 is a schematic diagram showing one embodiment of the present invention. Referring to Figure 4, a target assembly **10** is provided which includes a target vessel **12** and other components used to deliver materials to or from the target vessel **12** as will be described further below. The target vessel **12** is fabricated from a material having sufficient heat tolerance and sufficient structural strength to allow the target vessel **12** to maintain integrity without significant deformation during a process of bombardment of target material within the target vessel **12** by a cyclotron to produce a radioisotope ("the bombardment cycle"). The target vessel **12** defines a target chamber **14** adapted to hold a target material, such as heavy water. The target vessel **12** further defines a window on at least one side of the target chamber **14**. The window is covered with a thin sheet **17** of metal or similar particle-permeable material. The window and cover sheet **17** are adapted to allow high-energy particles

to enter the target chamber **14** and bombard a target material held within the target chamber **14** to produce at least one radioisotope. In several embodiments, the window and cover sheet **17** are positioned such that, when the target assembly **10** undergoes the bombardment cycle, the window and cover sheet **17** are most directly in the path of the particle stream within the cyclotron. In many embodiments of the present invention, the cover sheet **17** is welded to the body of the target vessel. The material forming the cover sheet **17** is selected to have physical properties permitting the through passage of charged particles (i.e. "particle-permeable") sufficient to allow at least a percentage of charged particles in the particle stream to pass through the cover sheet **17** to interact with the target material held within the target chamber **14**. In some embodiments, the cover sheet **17** is fabricated from a metal such as aluminum. In other embodiments, the cover sheet is fabricated from a material selected from the group consisting of Havar® and Arnavar. In several embodiments, the cover sheet **17** is secured to the target vessel **12** so as to ensure that liquid does not leak from the target chamber **14** through the window. In several embodiments, the cover sheet **17** is welded to the target vessel **12**. In several embodiments, the target assembly **10** is designed so that it can be used as an internal target with a cyclotron.

[0014] Referring to Figure 4, a target material input **1601** is provided in fluidic communication with the target chamber **14** to allow delivery of a target material to the target chamber **14**. A rinse water storage compartment **1609** is also provided in fluidic communication with the target chamber **14** to supply rinse water to the target chamber **14**. In the illustrated embodiment, the target chamber **14** is in fluidic communication with a load/unload tube **1605**, which is connected to a delivery valve **54**. The delivery valve **54** is, in turn, connected to a fill tube **1603** and a delivery tube **1607** and is configured to allow selective fluidic communication of the load/unload tube **1605** between the fill tube **1603** and the delivery tube **1607**. The delivery tube **1607** leads to a chemical production unit **40** as will be discussed further below. The fill tube **1603** is in fluidic communication with a water intake valve **52**.

The water intake valve **52** is, in turn, connected to the target material input **1601** and a rinse water storage compartment **1609**, and is configured to allow selective fluidic communication of the fill tube **1603** between the target material input **1601** and the rinse water storage compartment **1609**. A check valve **53** is positioned on the fill tube **1603** to ensure that water does not flow back through the fill tube **1603** toward the storage compartments.

[0015] A gas supply **30** is provided in communication with the target chamber **14** to keep the target material under pressure in the target chamber **14** during the bombardment process. In the illustrated embodiment, the target chamber **14** is in fluidic communication with a gas tube **30**, which is in turn in fluidic communication with a gas input tube **31** and a vent tube **32**. The gas input tube **31** is in fluidic communication with a gas supply **33** such as a gas storage container. In several embodiments, the gas supply **33** is configured to supply an inert gas, such as argon. The vent tube **32** is in fluidic communication with a gas output **34** such as an aperture leading to a gas storage unit or the open air. In certain embodiments, a filter **35** is provided to filter gas flowing through the vent tube **32** to the gas output **34**. In the illustrated embodiment, a gas output valve **51** is provided to regulate flow of gas through the vent tube **32** between the gas tube **30** and the gas output **34**, and similarly, an inert gas valve **55** is provided to regulate flow of gas through the gas input tube **31** between the gas supply **33** and the gas tube **30**.

[0016] During the bombardment cycle, first, the target chamber **14** is vented by opening the gas output valve **51**, which allows air to flow freely from target chamber **14** through the gas tube **30**, the open gas output valve **51**, the vent tube **32** and the filter **35** to the gas output **34**. Second, the delivery valve is adjusted to connect the load/unload tube **1605** with the fill tube **1603**; the water intake valve is adjusted to connect the fill tube **1603** with the target material storage compartment **1601**; and a preselected amount of heavy water or other target material is loaded into the target chamber **14** through the fill tube **1603** and the load/unload tube **1605**. Third, the

delivery valve **54** is closed and the target material in the target chamber **14** is placed under pressure by pumping high pressure argon or other inert gas into the target chamber **14** from the inert gas storage chamber **31**. Fourth, once the target material is under pressure, a high-energy particle beam from a cyclotron or other particle accelerator strikes the particle-permeable cover sheet **17** over the target chamber **14**. Some of the charged particles from the particle beam pass through the cover sheet **17** and interact with the target material in the target chamber **14**, producing the intended radioisotopes. After the bombardment with the particle beam has gone on for a pre-determined length of time, the bombardment ceases. In some embodiments, the target chamber **14** is then vented by closing the inert gas valve **55** and opening the gas output valve **51**. The gas output valve **51** is then closed.

[0017] Following the bombardment process, the transformed target material (with radioisotopes) is evacuated from the target chamber **14** and directed to a chemical processing unit **40**, where the radioisotopes formed within the target material may be combined with other reagents to synthesize a product such as a radiopharmaceutical. In this delivery cycle, the inert gas valve **55** is opened, the delivery valve **54** is adjusted to connect the load/unload tube **1605** with the delivery tube **1607**, and pressure from argon or other inert gas pushes the target material (with the radioisotopes) through the load/unload tube **1605** and the delivery tube **1607** to the chemical production unit **40**, where, in certain applications, the radioisotopes are reacted with other reagents to synthesize radiopharmaceuticals.

[0018] After the delivery cycle, if a rinse cycle is necessary, the target chamber **14** is first vented by opening the gas output valve **51**, which allows air to flow freely from target chamber **14** through the gas tube **30**, the open gas output valve **51**, the vent tube **32** and the filter **35** to the gas output **34**. Second, the water intake valve **52** is adjusted to allow a pre-selected amount of sterile rinse water to flow from the rinse water storage compartment **1609** through the fill tube **1603**; the delivery valve **54** is positioned to connect the fill tube **1603** and the load/unload tube **1605**; the rinse

water then flows through the load/unload tube **1605** into the target chamber **14**. Third, the rinse water is evacuated from the target chamber **14**: the gas output valve **51** is closed and the inert gas valve **55** is opened, allowing inert gas to flow from the inert gas storage container **33** through the gas input tube **31** and the gas tube **30**; inert gas under pressure is used to push the rinse water out of the target chamber **14** through the load/unload tube **1605**. Fourth, the delivery valve **54** is adjusted to connect the load/unload tube **1605** with the delivery tube **1607**; the rinse water is then pushed through the load/unload tube **1605** and the delivery tube **1607** to the chemical production unit **40**, where, in certain applications, the rinsed radioisotopes are reacted with other reagents to synthesize radiopharmaceuticals.

[0019] During the bombardment cycle, and particularly during the bombardment with the particle beam, the target vessel **12** absorbs high amounts of energy from the charged particles; most of this energy is converted into heat. Additionally, the target material, which is being excited by the charged particles and is under high pressure, also becomes heated and transfers some of its heat to the target vessel **12**. As shown in Figures 4 and 5, to keep the target vessel from overheating, a cooling system **20** is provided in thermal communication with the target vessel **12** to remove heat from the target vessel **12**, thereby keeping the target vessel **12** from overheating while the particle beam bombards the target chamber **14**. In the illustrated embodiment, the target vessel **12** is connected to a cooling water input **21** and a cooling water output **27** through a cooling water input tube **22** and a cooling water output tube **24**, respectively. In certain embodiments, at least one cross tube **23** is provided to connect the cooling water input tube **22** with the cooling water output tube **24**. The cooling system **20** is adapted to direct cool water from the cooling water input **21**, through the cooling water input tube **22** and through the cooling water output tube **24**, to the cooling water output **27** in order to cool the target vessel **12**. In some embodiments, water is used as the cooling medium, although other cooling substances are contemplated. Of course, those skilled in the art will recognize other configurations suitable for reducing thermal energy within the target vessel **12**.

during the bombardment process, and such configurations may be used without departing from the spirit and scope of the present invention.

[0020] Figures 6 through 11 illustrate another example embodiment of a target assembly according to the present invention. Figure 6 presents a perspective view of the target assembly **101**, and Figure 7 presents a close-up look at an exploded view of the target vessel **121**. Figures 8A and 8B show side views of the target assembly **101** from two additional perspectives, while Figures 9, 10, and 11 provide sectional views of the target vessel **121**.

[0021] As shown in Figure 6, the target assembly **101** includes four tubes (collectively, “the supply tubes”) that carry substances to and from the target vessel **121**. These supply tubes, in the illustrated example embodiment, include a load-unload tube **161** to deliver target material to the target chamber; a gas delivery tube **301** to supply inert gas to the target chamber; and a cooling water input tube **221** and cooling water output tube **241**, which together help to circulate water or other fluid through the target vessel **121** in order to keep the temperature of the target vessel **121** within pre-selected limits during the bombardment process.

[0022] The target assembly **101** further includes a support tube **450**. The smaller-diameter supply tubes **161**, **221**, **241**, **301** travel through the support tube **450** before connecting with the target vessel **121**. The supply tubes **161**, **221**, **241**, **301** and support tube **450** collectively comprise a support structure for holding the target vessel **121** in position and for delivering substances (including rinse water, inert gas, target material, and coolant) to the target vessel **121**. In some embodiments of the present invention, the target vessel **121** is welded to the support tube **450**. In some embodiments of the present invention, the support tube **450** tapers from a larger cross-section diameter to a smaller cross-section diameter at the point where the support tube **450** meets the target vessel **121**.

[0023] In some embodiments of the present invention, the target assembly 101 further includes a plug 460 that caps that end of the support tube 450 that is opposite the target vessel 121. As shown in Figure 6, the support tubes 161, 221, 241, 301 enter the support tube 450 through apertures in the plug 460, each tube passing through its respective aperture in a close frictional fit. The plug 460, along with the target vessel 121 welded to the other end of the support tube 460, creates a tight seal on the support tube 450, and in some embodiments the support tube 450 is vacuum sealed.

[0024] Each supply tube 161, 221, 241, 301 has at least one corresponding channel within the target vessel 121; generally each supply tube meets its corresponding channel at the surface where the target vessel 121 meets the support tube 450. Thus the load/unload tube 161 travels through the support tube 450 and meets with (and in some embodiments is welded to) a load/unload channel 161a within the target vessel 121, seen in Figures 9 and 11. The gas supply tube 301 meets a gas supply channel 301a within the target chamber, which gives way to a reflux chamber 303 proximate to the target chamber 141, as shown in the section view of Figure 11. The cooling water input tube 221 meets with an intra-target-vessel cooling water input tube 221a, which directs cooling water or fluid toward the bottom of the target vessel 121; there, as shown by the dashed arrows in Figure 11, the circulating cooling water or fluid fills the cooling water circulation chamber 241a, which is carved out of the target vessel 121; the water or fluid then exits the cooling water circulation chamber 241a and the target vessel 121 through the cooling water output tube 214, connected to the top of the target vessel 121.

[0025] It will be recognized by those with skill in the art that other configurations for a cooling water circulation system are possible and contemplated by this invention, and in particular it is to be noted that the target vessel in some embodiments comprises more than one water or fluid circulation channel.

[0026] In some embodiments of the present invention, the body of the target vessel 121 is fabricated from a single piece of material, such as stainless steel or another

metal. When the target vessel **121** begins as a single piece of metal, the various volumes within the target vessel **121**, such as the target chamber **141** or the load/unload channel **161a**, may be formed by drilling holes or cavities within the metal. In the illustrated example embodiment, as shown in the sectional views of Figures 9, 10, and 11, the target chamber **141** is carved out of the target vessel **121**, the vertical load/unload channel **161a** is drilled into the target vessel **121**, and a horizontal supplemental load/unload channel (or “cross-channel”) **163a** is drilled to connect the vertical load/unload channel **161a** with the target chamber **141**.

[0027] In some embodiments, the target chamber **141** is carved out of the target vessel **121** and then covered with the thin particle-permeable cover sheet **171**. In some embodiments of the present invention, the cover sheet **171** is fabricated from an alloy such as Havar® or Arnavar. In one particular embodiment, the cover sheet **171** consists of a Havar® sheet 0.5 mm thick. In some embodiments, the cover sheet **171** is then welded to the target vessel **121**. In some embodiments, such as the illustrated example embodiment in Figures 6 and 7, the cover sheet **171** is secured in place in front of the target chamber **141** between a gasket **174** and a front clamp **176**; as indicated in the exploded view in Figure 7, the front clamp **176** works in cooperation with a back clamp **178** to hold the gasket **174** and cover sheet **171** in place against the target chamber **141**, such that the cover sheet **171** covers the exposed side of the target chamber **141**. The front clamp **176** and the gasket **174** both contain apertures such that, in the area that is the target area of the particle beam during the bombardment process, the cover sheet **171** is the only solid material directly between the particle beam and the target chamber **141**. In some embodiments, such as the illustrated example embodiment shown in Figures 6 and 7, bolts or other fastening devices **191a-d** secure the front clamp **176** and the back clamp **178** together.

[0028] In some embodiments of the present invention, the target chamber **141** is coated with tantalum plating or a similar coating before being covered with the

cover sheet **171**. Tantalum plating helps to maintain the structural integrity of the target vessel **121** during the bombardment process; tantalum's high melting point and resistance to corrosion insulates the metal of the target vessel body **121** from the heated and volatile target material. It will be recognized by those with skill in the art that other configurations for a target vessel formed from a single piece of metal or other material are possible and contemplated by this invention.

[0029] Forming the body of the target vessel **121** from a single piece of material presents advantages over many target assemblies found in the prior art. The target vessel **121** formed from a single piece of material will prove more durable and enjoy a longer useful service life than a comparable target vessel that includes many different parts. Further, with fewer parts making up the target vessel, there is less chance of contamination from components such as the O-rings found in many prior art assemblies. The target vessel described herein also allows for the faster dissipation of residual radiation following the bombardment process. Moreover, the target vessel described herein, by omitting certain materials found in many prior art assemblies, when used does not result in the production of such undesired side products as Cobalt-68.

[0030] When the body of the target vessel **121** is fabricated from a single piece of material, it is necessary that the chosen material exhibit certain characteristics. The material must exhibit a tolerance for high heat from the bombardment process. The material must be able to withstand, without deformation that would compromise the integrity of the target vessel or interfere with the operation of the device, pressures of up to 100-250 psi, and possibly higher, from the inert gas used to pressurize the target material in the target chamber **141**. Further, the material must conduct heat well in order to transfer heat from the target chamber **141** to the cooling water circulation chamber **241a**, where circulating water or fluid is available to carry away excess heat. A thermal conductivity value of at least 12 W/(m*K) is recommended.

Stainless steel is one such material exhibiting these properties, but other materials are also contemplated.

[0031] The bombardment cycle for the target assembly **101** is similar to the bombardment cycle described above for the target assembly **10a**. The target chamber **141** is vented and the heavy water or other target material is loaded into the target chamber through the load/unload tube **161**, the load/unload channel **161a**, and the cross-channel **163a**. The contents of the target chamber **141** are placed under pressure by importing pressurized inert gas through the gas supply tube **301** and the gas supply channel **301a**. The target material is then altered by focusing a high-energy particle beam on the cover sheet **171** covering the target chamber **141**. As said above, the target material selected in many embodiments is heavy water. During this bombardment process, the bombardment of the heavy water in the target chamber **141** turns some of the heavy water into steam. This steam travels into the reflux chamber **303**, where, being out of the direct path of the particle beam and subject to cooling from the water circulation system and pressure from the pressurized gas, the steam condenses back into water.

[0032] As noted above, in some embodiments of the present invention, the cover sheet **171** is fabricated from an alloy such as Havar® or Arnavar. In one particular embodiment, the cover sheet **171** consists of a Havar® sheet 0.5 mm thick. In this embodiment, a proton beam strikes the Havar® cover sheet **171** with approximately 7.5 MeV energy; the cover sheet **171** allows 6.8 MeV to pass through to interact with the heavy water. In this embodiment, the heavy water is kept under approximately 100 to 250 psi of pressure from the inert gas pumped in through the gas supply tube **301** and gas supply channel **301a**. This pressure raises the boiling point of the heavy water and also helps to compress the target material within the target chamber **141**, thereby ensuring more interaction between the charged particles and the O-18 atoms, improving the yield of radioisotopes. Those of skill in the art will recognize that the particle-beam energies employed here, on the order of 7.5 Watts with a 1

micro-Amp current to produce a proton beam with 7.5 MeV energy, and the pressures involved, on the order of 100 to 250 psi, are considerably lower than the requirements for many systems in the prior art.

[0033] While the present invention has been illustrated by description of one embodiment, and while the illustrative embodiment has been described in detail, it is not the intention of the applicant to restrict or in any way limit the scope of the appended claims to such detail. Additional modifications will readily appear to those skilled in the art. The invention in its broader aspects is therefore not limited to the specific details, representative apparatus and methods, and illustrative examples shown and described. Accordingly, departures may be made from such details without departing from the spirit or scope of applicant's general inventive concept.

CLAIMS

What is claimed is:

1. A target assembly to produce a radioisotope from a target material comprising:
 - a target vessel having a body fabricated from a single piece of material, said target vessel defining a target chamber to hold the target material and to position the target material in the path of a beam of charged particles, whereby when the charged particles interact with the target material in said target chamber, at least one radioisotope is formed; and
 - a support structure to deliver target material to said target chamber, to pressurize the target material within said target chamber, to remove radioisotopes from said target chamber, and to cool said target vessel.
2. The target assembly of Claim 1 wherein said body of said target vessel is formed from a material capable of withstanding without compromising deformation pressures of up to 250 pounds per square inch.
3. The target assembly of Claim 1 wherein said body of said target vessel is formed from stainless steel.
4. The target assembly of Claim 1 wherein said body of said target vessel is formed from a material exhibiting thermal conductivity of at least 12 Watts per meter per Kelvin.
5. The target assembly of Claim 1 wherein said target chamber defines a window at least partially covered by a sheet of particle-permeable material, said

sheet being positioned to allow the beam of charged particles to penetrate the sheet to bombard the target material.

6. The target assembly of Claim 5 wherein said sheet of particle-permeable material is welded to said target vessel.

7. The target assembly of Claim 5 wherein said sheet of particle-permeable material is fabricated from a material selected from the group consisting of Havar, Arnavar, and aluminum.

8. The target assembly of Claim 7 wherein said sheet of particle-permeable material is secured to said target vessel by a clamp and gasket.

9. The target assembly of Claim 5 wherein said body of said target vessel is formed from stainless steel.

10. A target assembly to produce a radioisotope from a target material comprising:

a target vessel having a body fabricated from a single piece of material, said target vessel including a target chamber for holding the target material and for positioning the target material in the path of a beam of charged particles, whereby when the charged particles interact with the target material in said target chamber, radioisotopes are formed,

said target chamber being covered on at least one side by a sheet of material adapted to withstand the impact of a beam of charged particles, said sheet being positioned to cover an area directly in the path of the beam of charged particles, said sheet being adapted to permit the through passage of at least some charged particles, whereby when the beam of charged particles comes into contact with said sheet at least some charged particles in the beam of charged particles pass through said sheet to interact with the target material in said target chamber; and

a support structure for delivering target material to said target chamber, for pressurizing the target material within said target chamber, for removing radioisotopes from said target chamber, and for cooling said target vessel.

11. The target assembly of Claim 10 wherein said sheet is fabricated from Havar.

12. The target assembly of Claim 10 wherein said sheet is fabricated from Arnavar.

13. The target assembly of Claim 10 wherein said body of said target vessel is formed from stainless steel.

14. The target assembly of Claim 10 wherein said target vessel is fabricated from material capable of withstanding deformation pressures of up to 250 pounds per square inch.

15. The target assembly of Claim 10 wherein said target vessel is fabricated from material exhibiting a thermal conductivity of at least 12 Watts per meter per Kelvin.

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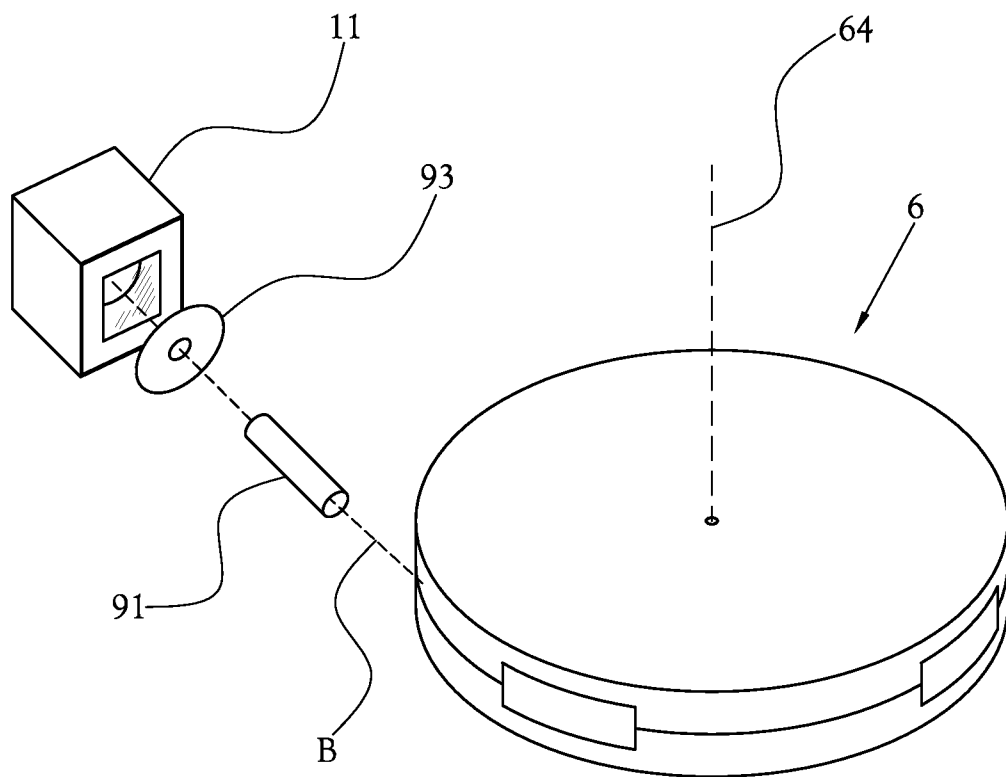


Fig.1
(PRIOR ART)

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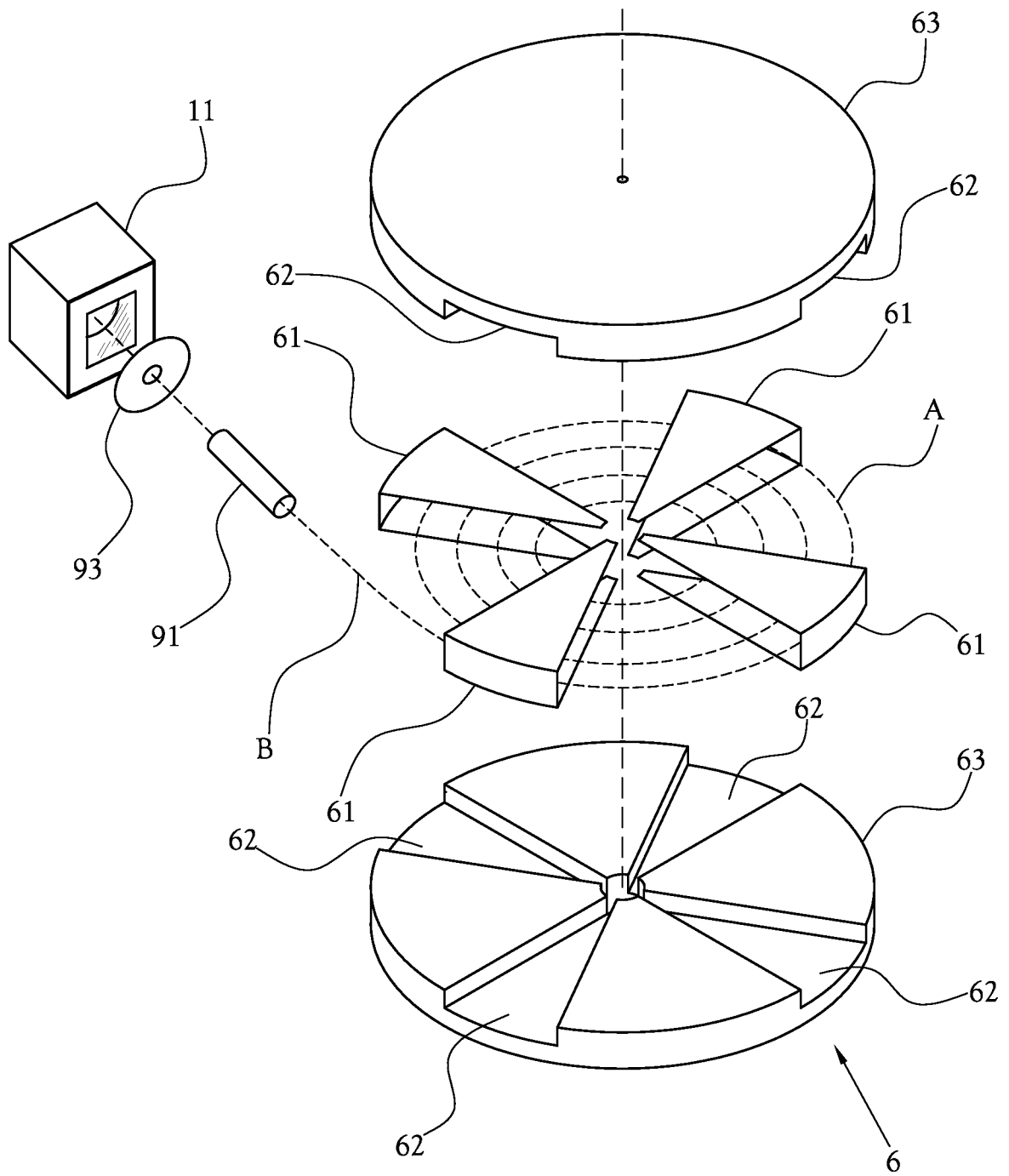


Fig.2

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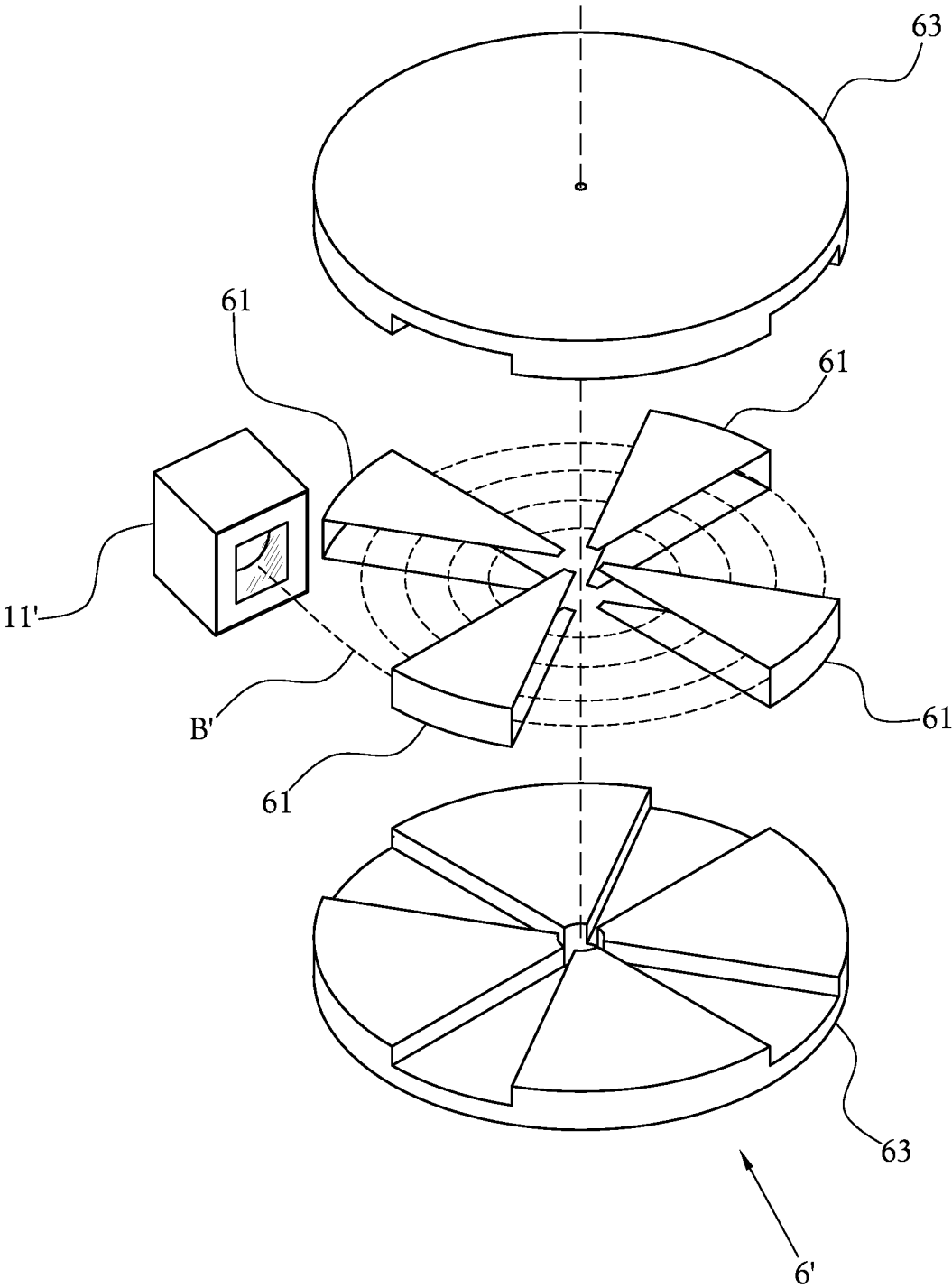
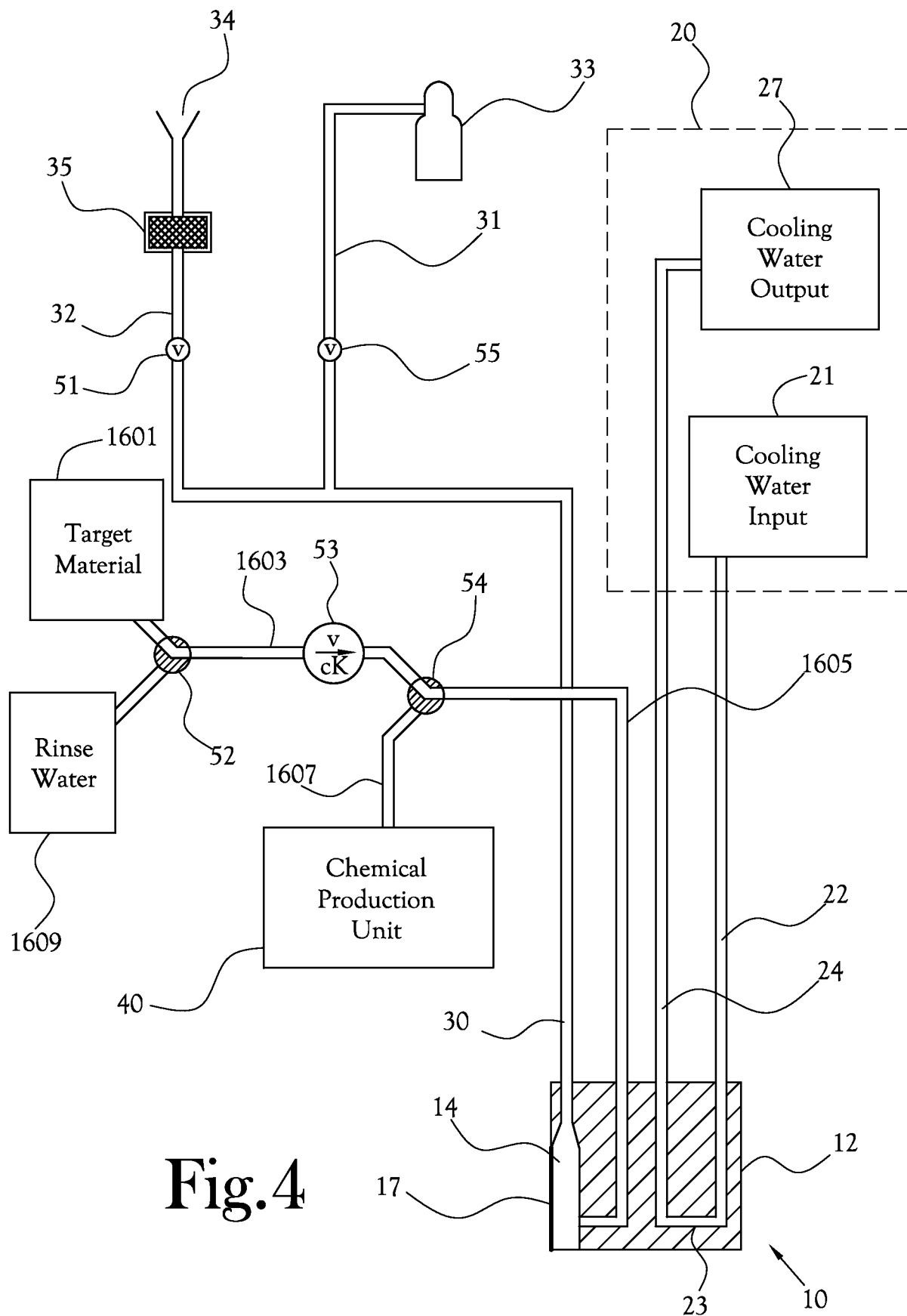


Fig.3

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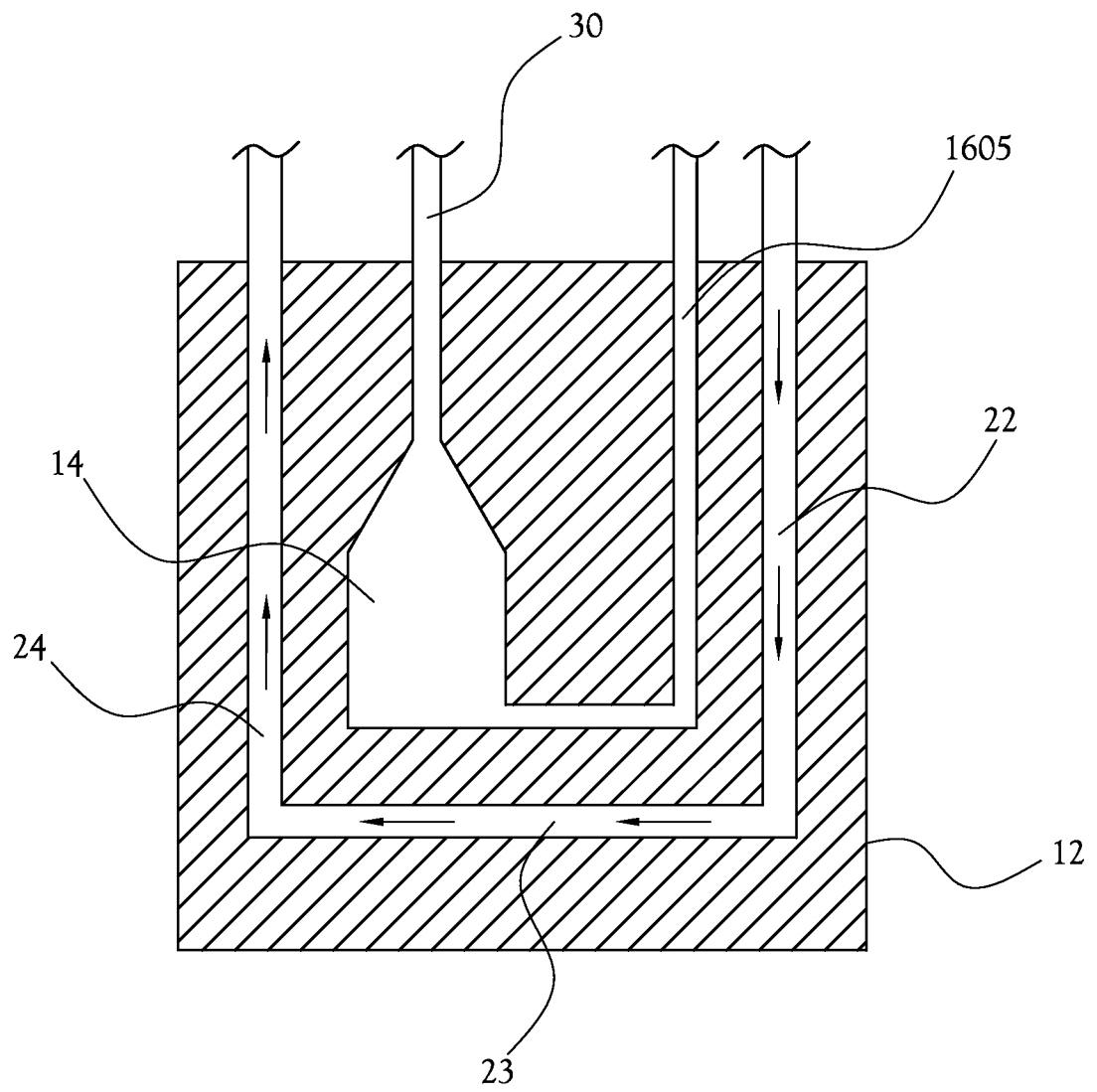


Fig.5

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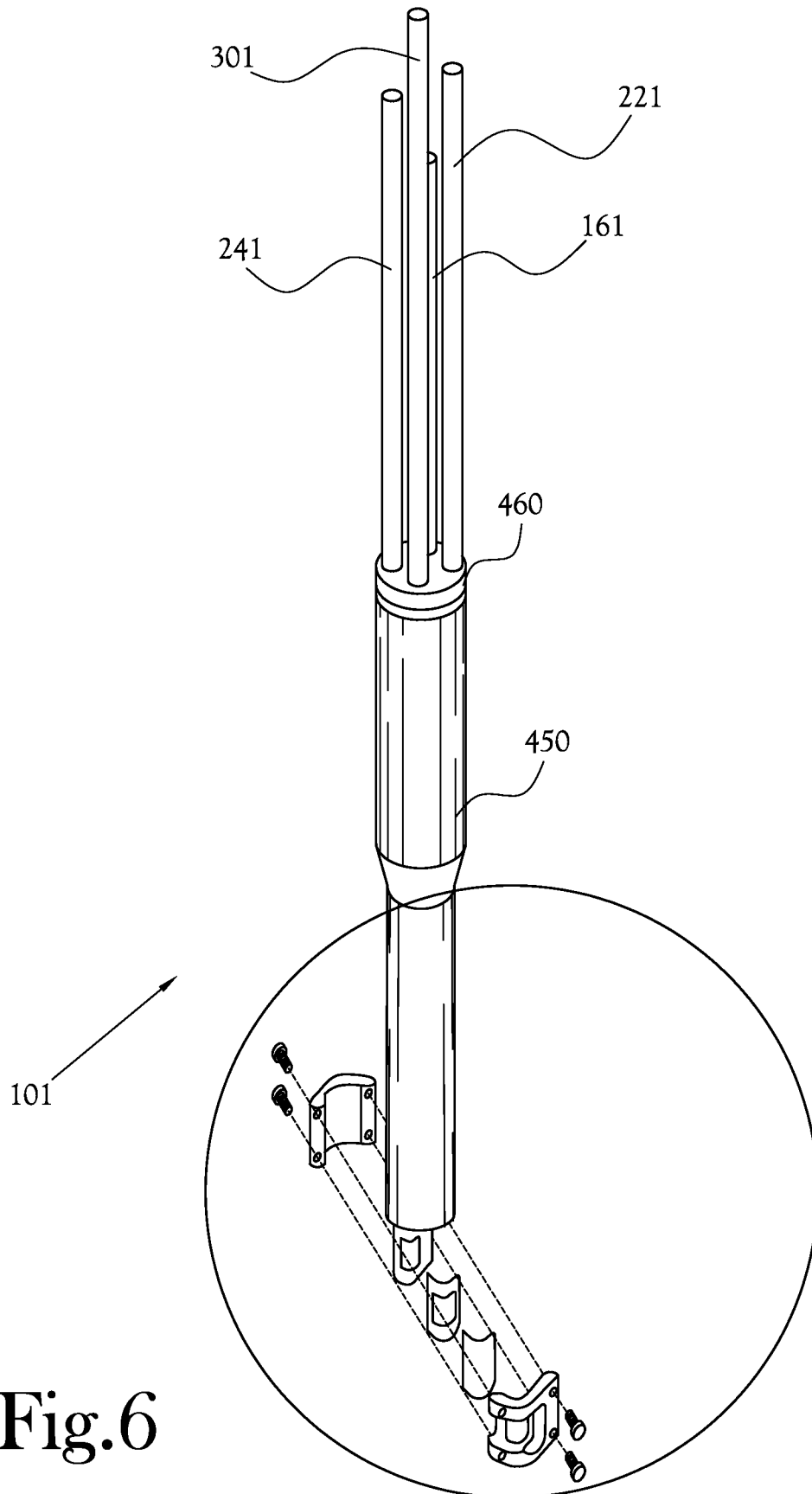
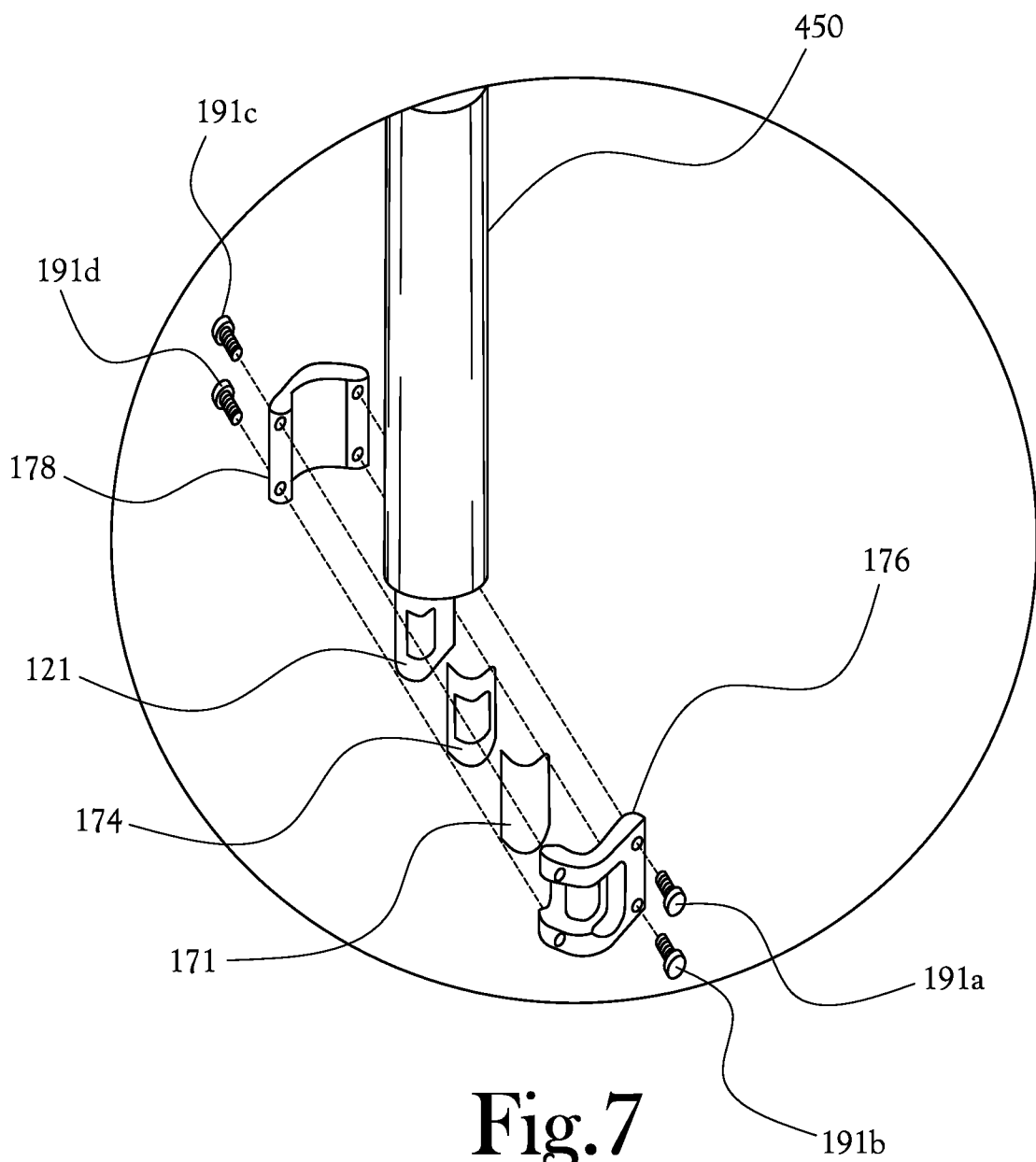


Fig.6

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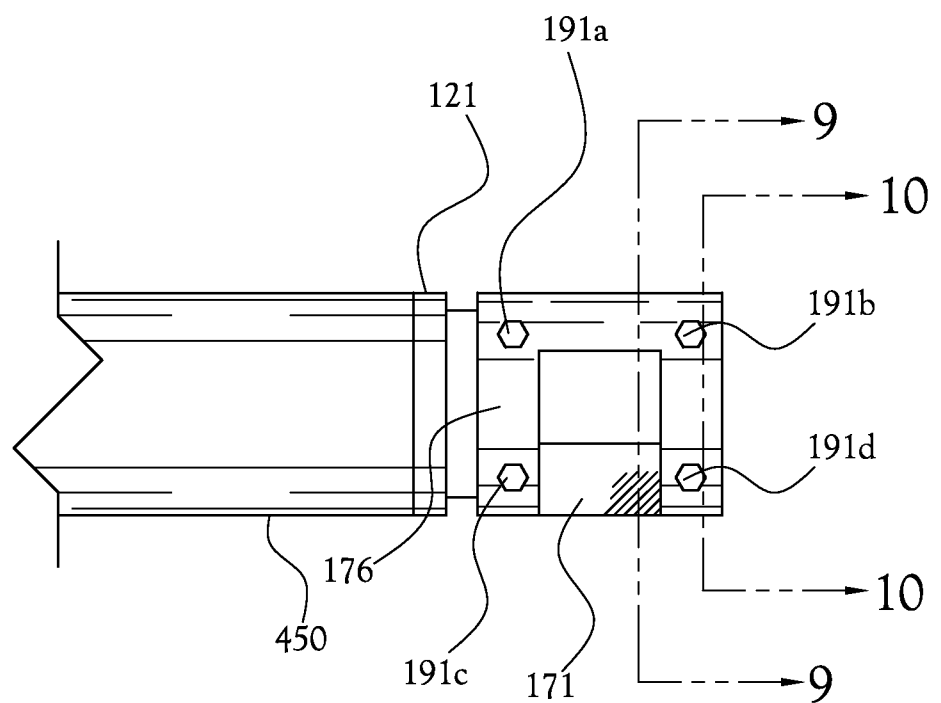
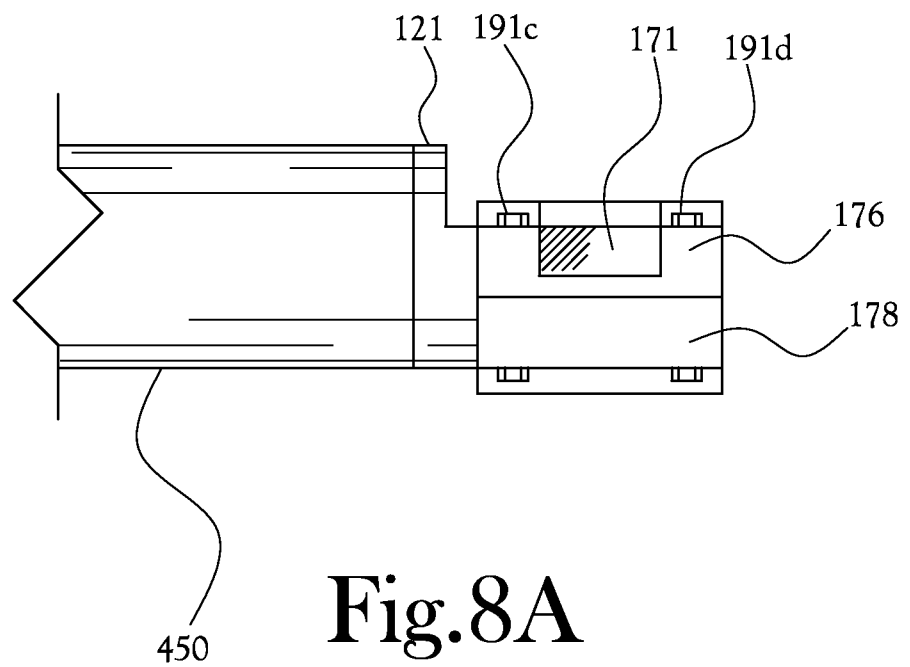


Fig.8B

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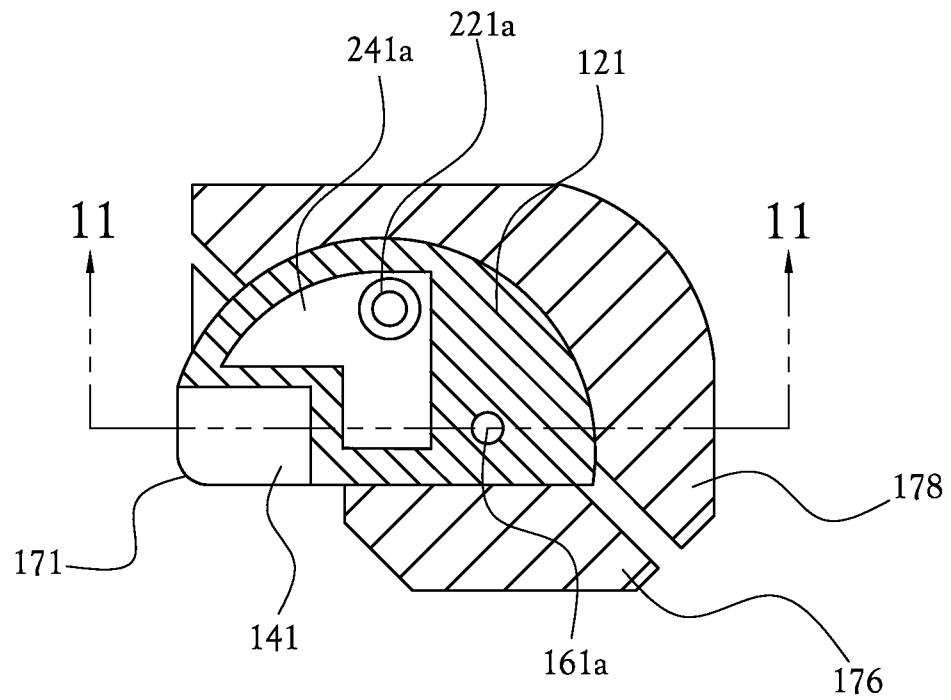


Fig.9

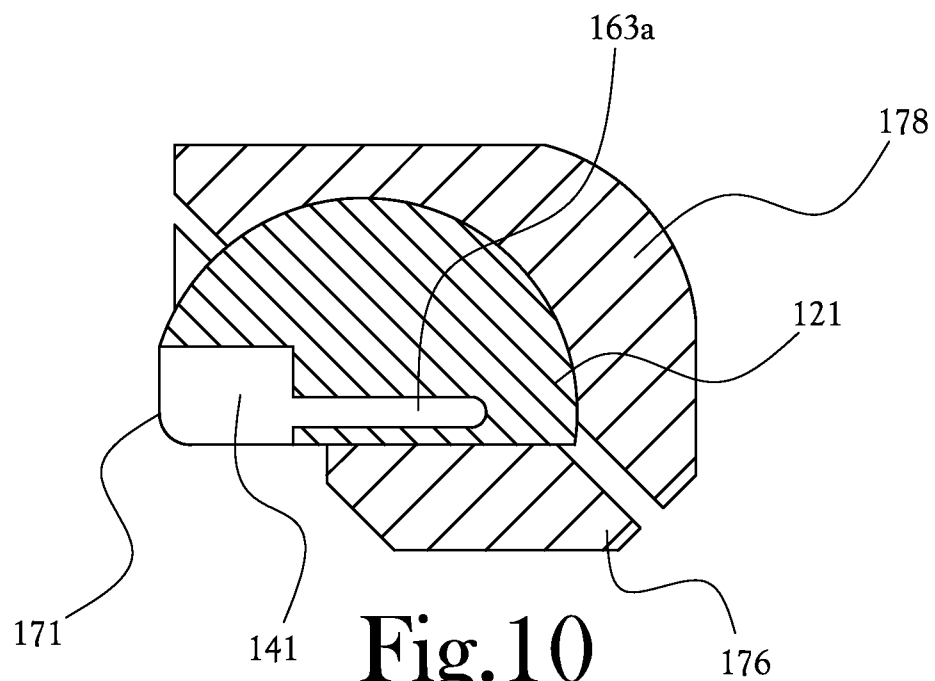


Fig.10

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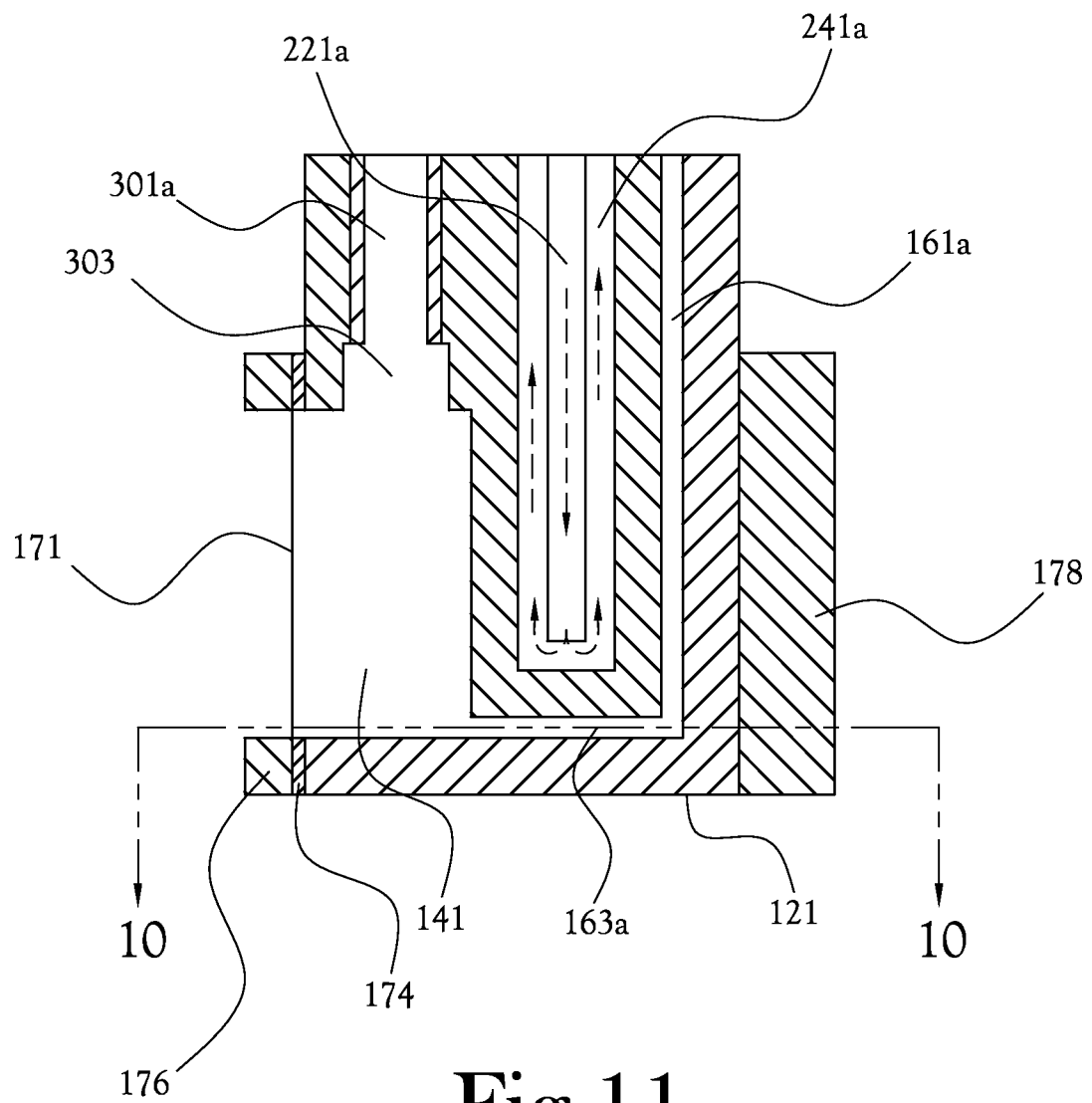


Fig.11

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US 12/58291

A. CLASSIFICATION OF SUBJECT MATTER

IPC(8) - G21G 1/10 (2012.01)

USPC - 376/190; 250/492.3

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

USPC: 376/190; 250/492.3

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched
 IPC(8): G21G 1/10 (2012.01); USPC: 376/190-202,156; 250/492.1,492.3,396R, 398
 (keyword limited; terms below)

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

PatBase (PatBase); PubWEST(USPT,PGPB,EPAB,JPAB); Google Scholar

Search Terms: radioisotope, target, vessel, pressure, steel, havar, thermal, conductivity, clamp, gasket, chamber, holder, amavar, aluminum, deformation, beam.

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X --- Y	US 6,011,825 A (WELCH et al.) 04 January 2000 (04.01.2000), entire document, especially; FIG 3A-3F, 4, col 1, ln 17-27, col 2, ln 57-65, col 9, ln 41 to col 12, ln 21, col 12, ln 41-62, col 14, ln 11-55, col 15, ln 32-57.	1 ----- 2-15
Y	US 2009/0090875 A1 (GELBART et al.) 09 April 2009 (09.04.2009), entire document, especially; para [0003], [0007], [0030], [0034], [0045], [0046].	2, 5-12, 13-15
Y	US 4,818,468 A (JUNGERMAN et al.) 04 April 1989 (04.04.1989), entire document, especially; col 1, ln 30-42, col 1, ln 61 to col 2, ln 4, col 6, ln 8-22.	3, 4, 9, 13, 15
Y	US 2005/0221358 A1 (CARRILLO et al.) 06 October 2005 (06.10.2005), entire document, especially; para [0499], [0589].	8
Y	Fosshag et al. "A target system for the production of 150 beams for ISAC." Published in: Nuclear Instruments and Methods in Physics Research A 480 (2002). Retrieved on 01 December 2012 (01.12.2012) from the Internet at: URL: http://144.206.159.178/fu/787/52266/917300.pdf . entire document, especially; pp 125, col 2.	12

☒ Further documents are listed in the continuation of Box C.

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"O" document referring to an oral disclosure, use, exhibition or other means

"P" document published prior to the international filing date but later than the priority date claimed

"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art

"&" document member of the same patent family

Date of the actual completion of the international search

26 November 2012 (26.11.2012)

Date of mailing of the international search report

11 DEC 2012

Name and mailing address of the ISA/US

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INTERNATIONAL SEARCH REPORT

International application No.

PCT/US 12/58291

C (Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	US 2006/0104401 A1 (JONGEN et al.) 18 May 2006 (18.05.2006), entire document, especially; FIG 1, para [0001], [0027]-[0030], [0058], [0059].	1-15
A	US 2010/0127188 A1 (NUTT) 27 May 2010 (27.05.2010), entire document, especially; FIG 2, 8, para [0017], [0052], [0054].	1-15
A	US 2005/0084055 A1 (ALVORD et al.) 21 April 2005 (21.04.2005), entire document, specially FIG 4-5, para [0004], [0032], [0033].	1-15