

(19) World Intellectual Property Organization
International Bureau



(43) International Publication Date
12 April 2007 (12.04.2007)

PCT

(10) International Publication Number
WO 2007/041031 A2

(51) International Patent Classification:

A61M 31/00 (2006.01)

(21) International Application Number:

PCT/US2006/037025

(22) International Filing Date:

21 September 2006 (21.09.2006)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

11/241,749 29 September 2005 (29.09.2005) US

(71) Applicant (for all designated States except US):

FLOWMEDICA, INC. [US/US]; 46563 Fremont Blvd., Fremont, CA 94538 (US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): **ELKINS, Jeffrey, M.** [US/US]; 12 Rebelo Lane, Novato, CA 94947 (US). **GOODSON, Harry, B.** [US/US]; 3596 Dickenson Common, Fremont, CA 94538 (US). **BALL, Craig, A.** [US/US]; 152 Belvedere Avenue, San Carlos, CA 94070

(US). **VALENCIA, Aurelio** [US/US]; 2249 Clark Avenue, East Palo Alto, CA 94303 (US). **PATEL, Samir, R.** [US/US]; 366 Sierra Vista Avenue, #6, Mountain View, CA 94043 (US). **HEKMAT, Neema** [US/US]; 725 Mariposa Avenue, #205, Mountain View, CA 94041 (US).

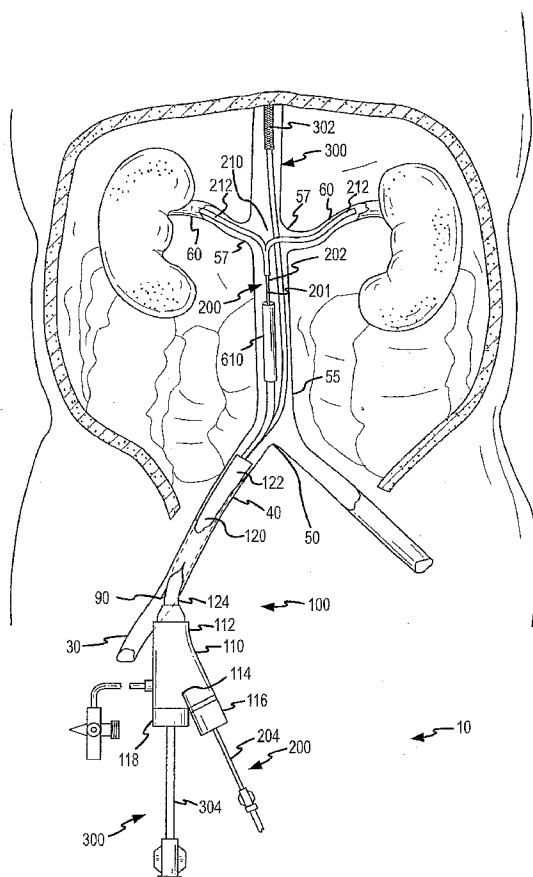
(74) Agents: **CASELL, Nathan** et al.; TOWNSEND AND TOWNSEND AND CREW LLP, Two Embarcadero Center, Eighth Floor, San Francisco, CA 94111-3834 (US).

(81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH,

[Continued on next page]

(54) Title: RENAL INFUSION SYSTEMS AND METHODS



(57) Abstract: Systems, devices, and methods for delivering treatment to the renal arteries are provided. Exemplary systems include a delivery catheter having a distal bifurcation, an introducer assembly comprising an introducer sheath in operative association with a Y-hub, wherein Y-hub includes a first port for receiving the delivery catheter and a second port for receiving a second catheter, and a constraint assembly for holding the distal bifurcation of the delivery catheter in a low-profile configuration when it is advanced distally beyond the introducer sheath.



GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

— *without international search report and to be republished upon receipt of that report*

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

RENAL INFUSION SYSTEMS AND METHODS

CROSS-REFERENCES TO RELATED APPLICATIONS

[0001] This application is a continuation-in-part of PCT Patent Application No.

- 5 PCT/US2003/29740 filed September 22, 2003 (Attorney Docket No. 022352-000600PC), which claims the benefit of U.S. Patent Application No. 10/251,915 filed September 20, 2002 (Attorney Docket No. 022352-000600US); and of PCT Patent Application No. PCT/US2004/008571 filed March 19, 2004 (Attorney Docket No. 022352-000900PC), which claims the benefit of U.S. Patent Application No. 60/508,751 filed October 2, 2003 (Attorney Docket No. 022352-000900US); and of U.S. Patent Application No. 11/084,738 filed March 16, 2005 (Attorney Docket No. 022352-001010US), which is a continuation-in-part of PCT Patent Application No. PCT/US2003/29744 filed September 22, 2003 (Attorney Docket No. 022352-001000PC), which claims the benefit of U.S. Patent Application No. 60/476,347 filed June 5, 2003 (Attorney Docket No. 022352-001000US) and of U.S. Patent Application No. 15 60/502,600 filed September 13, 2003 (Attorney Docket No. 022352-001400US); and of U.S. Patent Application No. 11/084,434 filed March 18, 2005 (Attorney Docket No. 022352-001110US), which is a continuation of PCT Patent Application No. PCT/2003/29995 filed September 22, 2003 (Attorney Docket No. 022352-001100PC), which claims the benefit of U.S. Patent Application No. 60/412,343 filed September 20, 2002 (Attorney Docket No. 20 022352-000700US), and of U.S. Patent Application No. 60/412,476 filed September 20, 2002 (Attorney Docket No. 022352-000800US), and of U.S. Patent Application No. 60/479,329 filed June 17, 2003 (Attorney Docket No. 022352-001100US), and of U.S. Patent Application No. 60/502,389 filed September 13, 2003 (Attorney Docket No. 022352-001500US); and of U.S. Patent Application No. 11/083,802 filed March 18, 2005 (Attorney Docket No. 022352-001210US), which is a continuation of PCT Patent Application No. PCT/US2003/29743 filed September 22, 2003 (Attorney Docket No. 022352-001200PC), which claims the benefit of U.S. Patent Application No. 60/486,349 filed July 10, 2003 (Attorney Docket No. 022352-001200US); and of U.S. Patent Application No. 11/084,295 filed March 18, 2005 (Attorney Docket No. 022352-001310US), which is a continuation of PCT Patent Application No. PCT/US2003/29585 filed September 22, 2003 (Attorney Docket No. 022352-001300PC), which claims the benefit of U.S. Patent Application No. 60/486,206 filed July 9, 2003 (Attorney Docket No. 022352-001300US) and of U.S. Patent Application

No. 60/502,399 filed September 13, 2003 (Attorney Docket No. 022352-001600US); and of PCT Patent Application No. PCT/US2004/008573 filed March 19, 2004 (Attorney Docket No. 022352-001900PC), which claims the benefit of U.S. Patent Application No. 60/505,281 filed September 22, 2003 (Attorney Docket No. 022352-001700US) and of U.S. Patent Application No. 60/543,671 filed February 9, 2004 (Attorney Docket No. 022352-001900US); and of U.S. Patent Application No. 11/073,421 filed March 4, 2005 (Attorney Docket No. 022352-002020US), which claims the benefit of U.S. Patent Application No. 60/550,632 filed March 4, 2004 (Attorney Docket No. 022352-002000US) and of U.S. Patent Application No. 60/550,774 filed March 5, 2004 (Attorney Docket No. 022352-002010US); and of U.S. Patent Application No. 11/129,101 filed May 13, 2005 (Attorney Docket No. 022352-002120US), which claims the benefit of U.S. Patent Application No. 60/571,057 filed May 14, 2004 (Attorney Docket No. 022352-002100US) and of U.S. Patent Application No. 60/612,731 filed September 24, 2004 (Attorney Docket No. 022352-002110US); and claims the benefit of U.S. Provisional Patent Application No. 60/612,801 filed September 24, 2004 (Attorney Docket No. 022352-002700US). This application is also related to U.S. Patent No. 6,749,598 filed January 11, 1999 (Attorney Docket No. 022352-000300US), U.S. Patent Application No. 09/562,493 filed May 1, 2000 (Attorney Docket No. 022352-000400US), U.S. Patent Application No. 09/724,691 filed November 28, 2000 (Attorney Docket No. 022352-000500US), PCT Patent Application No. PCT/US2003/29586 filed September 22, 2003 (Attorney Docket No. 022352-001800US), U.S. Patent Application No. 60/582,870 filed June 24, 2004 (Attorney Docket No. 022352-002200US), and U.S. Patent Application No. 11/167,056 filed June 23, 2005 (Attorney Docket No. 022352-002310US). The entire contents of each of these applications and their priority filings is incorporated herein by reference for all purposes.

BACKGROUND OF THE INVENTION

[0002] This invention pertains generally to medical device systems and methods for delivering treatment to internal body lumens. More specifically, it is related to intra aortic renal treatment delivery systems and methods.

[0003] Many different medical device systems and methods have been previously described for locally delivering fluids or other agents into various body regions, including body lumens such as vessels, or other body spaces such as organs or heart chambers. Local fluid delivery systems may include drugs or other agents, or may even include locally

delivering the body's own fluids, such as artificially enhanced blood transport, for example either entirely within the body such as directing or shunting blood from one place to another, or in extracorporeal modes such as via external blood pumps and the like. Local agent delivery systems are herein generally intended to relate to introduction of a foreign

5 composition as an agent into the body, which may include drugs or other useful or active agents, and may be in a fluid form or other form such as gels, solids, powders, gases, and the like. It is to be understood that reference to only one of the terms fluid, drug, or agent with respect to local delivery descriptions may be made variously in this disclosure for illustrative purposes, but is not generally intended to be exclusive or omissive of the others; they are to
10 be considered interchangeable where appropriate according to one of ordinary skill unless specifically described to be otherwise.

[0004] In general, local agent delivery systems and methods are often used for the benefit of achieving relatively high, localized concentrations of agent where injected within the body in order to maximize the intended effects there and while minimizing unintended peripheral
15 effects of the agent elsewhere in the body. Where a particular dose of a locally delivered agent may be efficacious for an intended local effect, the same dose systemically delivered would be substantially diluted throughout the body before reaching the same location. The agent's intended local effect is equally diluted and efficacy is compromised. Thus systemic agent delivery requires higher dosing to achieve the required localized dose for efficacy,
20 often resulting in compromised safety due to for example systemic reactions or side effects of the agent as it is delivered and processed elsewhere throughout the body other than at the intended target.

[0005] Various diagnostic systems and procedures have been developed using local delivery of dye (e.g. radiopaque contrast agent) or other diagnostic agents, wherein an
25 external monitoring system is able to gather important physiological information based upon the diagnostic agent's movement or assimilation in the body at the location of delivery and/or at other locations affected by the delivery site. Angiography is one such practice using a hollow, tubular angiography catheter for locally injecting radiopaque dye into a blood chamber or vessel, such as for example coronary arteries in the case of coronary angiography,
30 or in a ventricle in the case of cardiac ventriculography.

[0006] Other systems and methods have been reported for locally delivering therapeutic agent into a particular body tissue within a patient via a body lumen. For example,

angiographic catheters of the type just described above, and other similar tubular delivery catheters, have also been reported for use in locally injecting treatment agents through their delivery lumens into such body spaces within the body. More detailed examples of this type include local delivery of thrombolytic drugs such as TPATM, heparin, cumadin, or urokinase into areas of existing clot or thrombogenic implants or vascular injury. In addition, various balloon catheter systems have also been reported for local administration of therapeutic agents into target body lumens or spaces, and in particular associated with blood vessels. More specific previously disclosed of this type include balloons with porous or perforated walls that elute drug agents through the balloon wall and into surrounding tissue such as blood vessel walls. Yet further examples for localized delivery of therapeutic agents include various multiple balloon catheters that have spaced balloons that are inflated to engage a lumen or vessel wall in order to isolate the intermediate catheter region from in-flow or out-flow across the balloons. According to these examples, a fluid agent delivery system is often coupled to this intermediate region in order to fill the region with agent such as drug that provides an intended effect at the isolated region between the balloons.

[0007] The diagnosis or treatment of many different types of medical conditions associated with various different systems, organs, and tissues, may also benefit from the ability to locally deliver fluids or agents in a controlled manner. In particular, various conditions related to the renal system would benefit a great deal from an ability to locally deliver of therapeutic, prophylactic, or diagnostic agents into the renal arteries.

[0008] Acute renal failure ("ARF") is an abrupt decrease in the kidney's ability to excrete waste from a patient's blood. This change in kidney function may be attributable to many causes. A traumatic event, such as hemorrhage, gastrointestinal fluid loss, or renal fluid loss without proper fluid replacement may cause the patient to go into ARF. Patients may also become vulnerable to ARF after receiving anesthesia, surgery, or a-adrenergic agonists because of related systemic or renal vasoconstriction. Additionally, systemic vasodilation caused by anaphylaxis, and anti-hypertensive drugs, sepsis or drug overdose may also cause ARF because the body's natural defense is to shut down, i.e., vasoconstrict, non-essential organs such as the kidneys. Reduced cardiac output caused by cardiogenic shock, congestive heart failure, pericardial tamponade, or massive pulmonary embolism creates an excess of fluid in the body, which can exacerbate congestive heart failure. For example, a reduction in blood flow and blood pressure in the kidneys due to reduced cardiac output can in turn result

in the retention of excess fluid in the patient's body, leading, for example, to pulmonary and systemic edema.

[0009] Previously known methods of treating ARF, or of treating acute renal insufficiency associated with congestive heart failure ("CHF"), involve administering drugs. Examples of such drugs that have been used for this purpose include, without limitation: vasodilators, including for example papavarine, fenoldopam mesylate, calcium-channel blockers, atrial natriuretic peptide (ANP), acetylcholine, nifedipine, nitroglycerine, nitroprusside, adenosine, dopamine, and theophylline; antioxidants, such as for example acetylcysteine; and diuretics, such as for example mannitol, or furosemide. However, many of these drugs, when administered in systemic doses, have undesirable side effects. Additionally, many of these drugs would not be helpful in treating other causes of ARF. A septic shock patient with profound systemic vasodilation often has concomitant severe renal vasoconstriction, however administering vasodilators to dilate the renal artery to a patient suffering from systemic vasodilation would compound the vasodilation system wide. In addition, for patients with severe CHF (e.g., those awaiting heart transplant), mechanical methods, such as hemodialysis or left ventricular assist devices, may be implemented. Surgical device interventions, such as hemodialysis, however, generally have not been observed to be highly efficacious for long-term management of CHF. Such interventions would also not be appropriate for many patients with strong hearts suffering from ARF.

[0010] The renal system in many patients may also suffer from a particular fragility, or otherwise general exposure, to potentially harmful effects of other medical device interventions. For example, the kidneys as one of the body's main blood filtering tools may suffer damage from exposed to high density radiopaque contrast dye, such as during coronary, cardiac, or neuro angiography procedures. One particularly harmful condition known as "radiocontrast nephropathy" or "RCN" is often observed during such procedures, wherein an acute impairment of renal function follows exposure to such radiographic contrast materials, typically resulting in a rise in serum creatinine levels of more than 25% above baseline, or an absolute rise of 0.5 mg/dl within 48 hours. Therefore, in addition to CHF, renal damage associated with RCN is also a frequently observed cause of ARF. In addition, the kidneys' function is directly related to cardiac output and related blood pressure into the renal system. These physiological parameters, as in the case of CHF, may also be significantly compromised during a surgical intervention such as an angioplasty, coronary artery bypass, valve repair or replacement, or other cardiac interventional procedure.

Therefore, the various drugs used to treat patients experiencing ARF associated with other conditions such as CHF have also been used to treat patients afflicted with ARF as a result of RCN. Such drugs would also provide substantial benefit for treating or preventing ARF associated with acutely compromised hemodynamics to the renal system, such as during surgical interventions.

[0011] There would be great advantage therefore from an ability to locally deliver such drugs into the renal arteries, in particular when delivered contemporaneous with surgical interventions, and in particular contemporaneous with radiocontrast dye delivery. However, many such procedures are done with medical device systems, such as using guiding catheters or angiography catheters having outer dimensions typically ranging between about 4 French to about 12 French, and ranging generally between about 6 French to about 8 French in the case of guide catheter systems for delivering angioplasty or stent devices into the coronary or neurovascular arteries (e.g. carotid arteries). These devices also are most typically delivered to their respective locations for use (e.g. coronary ostia) via a percutaneous, transluminal access in the femoral arteries and retrograde delivery upstream along the aorta past the region of the renal artery ostia. A Seldinger access technique to the femoral artery involves relatively controlled dilation of a puncture hole to minimize the size of the intruding window through the artery wall, and is a preferred method where the profiles of such delivery systems are sufficiently small. Otherwise, for larger systems a "cut-down" technique is used involving a larger, surgically made access window through the artery wall.

[0012] Accordingly, an intra aortic renal agent delivery system for contemporaneous use with other retrogradely delivered medical device systems, such as of the types just described above, would preferably be adapted to allow for such interventional device systems, in particular of the types and dimensions just described, to pass upstream across the renal artery ostia (a) while the agent is being delivered into the renal arteries, and (b) while allowing blood to flow downstream across the renal artery ostia, and (c) in an overall cooperating system that allows for Seldinger femoral artery access. Each one of these features (a), (b), or (c), or any sub-combination thereof, would provide significant value to patient treatment; an intra aortic renal delivery system providing for the combination of all three features is so much the more valuable.

[0013] Notwithstanding the clear needs for and benefits that would be gained from such intra aortic drug delivery into the renal system, the ability to do so presents unique challenges as follows.

5 [0014] In one regard, the renal arteries extend from respective ostia along the abdominal aorta that are significantly spaced apart from each other circumferentially around the relatively very large aorta. Often, these renal artery ostia are also spaced from each other longitudinally along the aorta with relative superior and inferior locations. This presents a unique challenge to deliver drugs or other agents into the renal system on the whole, which requires both kidneys to be fed through these separate respective arteries via their uniquely
10 positioned and substantially spaced apart ostia. This becomes particularly important where both kidneys may be equally at risk, or are equally compromised, during an invasive upstream procedure – or, of course, for any other indication where both kidneys require renal drug delivery. Thus, an appropriate intra aortic delivery system for such indications would preferably be adapted to feed multiple renal arteries perfusing both kidneys.

15 [0015] In another regard, mere delivery of an agent into the natural, physiologic blood flow path of the aorta upstream of the kidneys may provide some beneficial, localized renal delivery versus other systemic delivery methods, but various undesirable results still arise. In particular, the high flow aorta immediately washes much of the delivered agent beyond the intended renal artery ostia. This reduces the amount of agent actually perfusing the renal
20 arteries with reduced efficacy, and thus also produces unwanted loss of the agent into other organs and tissues in the systemic circulation (with highest concentrations directly flowing into downstream circulation).

[0016] In still a further regard, various known types of tubular local delivery catheters, such as angiographic catheters, other “end-hole” catheters, or otherwise, may be positioned with
25 their distal agent perfusion ports located within the renal arteries themselves for delivering agents there, such as via a percutaneous transluminal procedure via the femoral arteries, or from other access points such as brachial arteries and the like. However, such a technique may also provide less than completely desirable results.

[0017] For example, such seating of the delivery catheter distal tip within a renal artery
30 may be difficult to achieve from within the large diameter/high flow aorta, and may produce harmful intimal injury within the artery. Also, where multiple kidneys must be infused with agent, multiple renal arteries must be cannulated, either sequentially with a single delivery

device, or simultaneously with multiple devices. This can become unnecessarily complicated and time consuming and further compound the risk of unwanted injury from the required catheter manipulation. Moreover, multiple dye injections may be required in order to locate the renal ostia for such catheter positioning, increasing the risks associated with contrast agents on kidney function (e.g. RCN) – the very organ system to be protected by the agent delivery system in the first place. Still further, the renal arteries themselves, possibly including their ostia, may have pre-existing conditions that either prevent the ability to provide the required catheter seating, or that increase the risks associated with such mechanical intrusion. For example, the artery wall may be diseased or stenotic, such as due to atherosclerotic plaque, clot, dissection, or other injury or condition. Finally, among other additional considerations, previous reports have yet to describe an efficacious and safe system and method for positioning these types of local agent delivery devices at the renal arteries through a common introducer or guide sheath shared with additional medical devices used for upstream interventions, such as angiography or guide catheters. In particular, to do so concurrently with multiple delivery catheters for simultaneous infusion of multiple renal arteries would further require a guide sheath of such significant dimensions that the preferred Seldinger vascular access technique would likely not be available, instead requiring the less desirable “cut-down” technique.

[0018] In addition to the various needs for delivering agents into branch arteries described above, much benefit may also be gained from simply enhancing blood perfusion into such branches, such as by increasing the blood pressure at their ostia. In particular, such enhancement would improve a number of medical conditions related to insufficient physiological perfusion into branch vessels, and in particular from an aorta and into its branch vessels such as the renal arteries.

[0019] Certain previous reports have provided surgical device assemblies and methods intended to enhance blood delivery into branch arteries extending from an aorta. For example, intra-aortic balloon pumps (IABPs) have been disclosed for use in diverting blood flow into certain branch arteries. One such technique involves placing an IABP in the abdominal aorta so that the balloon is situated slightly below (proximal to) the branch arteries. The balloon is selectively inflated and deflated in a counterpulsation mode (by reference to the physiologic pressure cycle) so that increased pressure distal to the balloon directs a greater portion of blood flow into principally the branch arteries in the region of their ostia. However, the flow to lower extremities downstream from such balloon system

can be severely occluded during portions of this counterpulsing cycle. Moreover, such previously reported systems generally lack the ability to deliver drug or agent to the branch arteries while allowing continuous and substantial downstream perfusion sufficient to prevent unwanted ischemia.

5 [0020] It is further noted that, despite the renal risks described in relation to radiocontrast dye delivery, and in particular RCN, in certain circumstances delivery of such dye or other diagnostic agents is indicated specifically for diagnosing the renal arteries themselves. For example, diagnosis and treatment of renal stenosis, such as due to atherosclerosis or dissection, may require dye injection into a subject renal artery. In such circumstances,
10 enhancing the localization of the dye into the renal arteries may also be desirable. In one regard, without such localization larger volumes of dye may be required, and the dye lost into the downstream aortic flow may still be additive to impacting the kidney(s) as it circulates back there through the system. In another regard, an ability to locally deliver such dye into the renal artery from within the artery itself, such as by seating an angiography catheter there,
15 may also be hindered by the same stenotic condition requiring the dye injection in the first place (as introduced above). Still further, patients may have stent-grafts that may prevent delivery catheter seating.

[0021] Notwithstanding the interest and advances toward delivering agents for treatment or diagnosis of organs or tissues, the previously reported systems and methods summarized
20 immediately above often lack the ability to effectively deliver agents from within a main artery and locally into substantially only branch arteries extending therefrom while allowing the passage of substantial blood flow and/or other medical devices through the main artery past the branches. This is in particular the case with previously reported renal treatment and diagnostic devices and methods, which do not adequately provide for local delivery of agents
25 into the renal system from a location within the aorta while allowing substantial blood flow continuously downstream past the renal ostia and/or while allowing distal medical device assemblies to be passed retrogradely across the renal ostia for upstream use. Much benefit would be gained if agents, such as protective or therapeutic drugs or radiopaque contrast dye, could be delivered to one or both of the renal arteries in such a manner.

30 [0022] Several more recently reported advances have included local flow assemblies using tubular members of varied diameters that divide flow within an aorta adjacent to renal artery ostia into outer and inner flow paths substantially perfusing the renal artery ostia and

downstream circulation, respectively. Such reports further include delivering fluid agent primarily into the outer flow path for substantially localized delivery into the renal artery ostia. These reported systems and methods represent exciting new developments toward localized diagnosis and treatment of pre-existing conditions associated with branch vessels from main vessels in general, and with respect to renal arteries extending from abdominal aortas in particular.

[0023] However, such previously reported designs would still benefit from further modifications and improvements in order to: maximize mixing of a fluid agent within the entire circumference of the exterior flow path surrounding the tubular flow divider and perfusing multiple renal artery ostia; use the systems and methods for prophylaxis and protection of the renal system from harm, in particular during upstream interventional procedures; maximize the range of useful sizing for specific devices to accommodate a wide range of anatomic dimensions between patients; and optimize the construction, design, and inter-cooperation between system components for efficient, atraumatic use.

[0024] A need still exists for improved devices and methods that allow the physician to leave an introducer sheath in place while advancing and retracting a jacket sheath. Similarly, a need still exists for improved devices and methods that provide a short introducer sheath that allows for a one-size-fits-all approach for placing a delivery catheter in the patient's body. Relatedly, the need still exists for improved approaches that do not require an introducer sheath that is optimally sized in each case according to the patient's anatomy (e.g. the distance from the puncture site to the renal artery origins or ostia). What is more, the need still exists for improved devices and methods that avoid the need to have excess external sheath length extending proximally from the entry site, as may be the case where the distance between the puncture site and the deployment site is significantly less the length of a standard introducer sheath. Further, a need still exists for improved devices and methods that create less interference or friction when the physician maneuvers various components of the catheter device. A need still exists for improved approaches that avoid or minimize the risk of having an inadequate length for an auxiliary catheter such as a coronary catheter to reach its intended site.

[0025] A need still exists for improved devices and methods that can eliminate many of the difficulties associated with managing iliac, aortic, and other vascular tortuosity. Further, a need still exists for improved approaches that minimize the potential for clot formation within

an introducer sheath and a Y-hub. Relatedly, a need still exists for improved approaches that decrease the risk of clots dislodging when adjunctive commercial products are passed through the introducer sheath, particularly at advancement. And the need still exists for improved approaches that provide a full-covered advancement of bifurcated catheter to the intended deployment site. Further, the need still exists for improved devices and methods that facilitate adjunctive coronary procedures, as well as various peripheral procedures. The present invention provides solutions for at least some of such needs.

BRIEF SUMMARY OF THE INVENTION

[0026] The present invention provides a treatment delivery system that is easy to use and minimizes the time required for many surgical procedures. With the incorporation of an improved introducer sheath, an integrated introducer and catheter system can provide universal application for a broad spectrum of patients, for various types of catheterization protocols, regardless of the unique anatomical features of the individual patient. Moreover, a moveable cover jacket sheath or other constraining assembly or structure allows for advancement and retraction of distal catheter tips or extensions in a collapsed or captured state. Constraining assemblies may include jackets, sheaths, loops, lassos, rings, and the like. The present invention also provides for improved guide wire guidance and advancement of the cover jacket sheath and catheter assembly for advancement. Guide wire systems can be of a coaxial design (e.g wire inside cover jacket sheath) or of a monorail type design, including a split-tube type of monorail to allow the rail to snap onto and off of the catheter shaft. Further, the present invention provides several improved cover jacket sheath configurations, which can allow blood passage therethrough, to avoid stagnant blood and potential thrombus formation within the jacket sheath. Many jacket sheath configurations may be sealed during the dwell period, including those which allow infusate to fill the sealed inner jacket space volume to prevent thrombus formation. It is further appreciated that the present invention provides a highly integrated catheter and sheath system where many components, including the constraining jacket sheath, remain together without extraneous tubes, wires, and the like protruding outside the introducer assembly's Y-hub during the dwell period.

[0027] These present embodiments therefore are particularly useful in intra aortic renal drug delivery systems introduces from a position proximal to the renal arteries; however, it is contemplated that these systems and methods may be suitably modified for use in other

anatomical regions and for other medical conditions without departing from the broad scope of various of the aspects illustrated by the embodiments. For example, intra aortic fluid delivery according to various of these embodiments benefits from particular dimensions, shapes, and constructions for the subject devices herein described. However, suitable
5 modifications may be made to deliver fluids to other multi-lateral branch structures from main body spaces or lumens such as other locations within the vasculature, including the right and left coronary artery ostia, fallopian tubes stemming from a uterus, or the gastrointestinal tract.

[0028] In a first aspect, the present invention provides a method for positioning a delivery
10 catheter in the renal arteries. The method can include positioning an introducer sheath in an iliac artery, advancing a renal delivery catheter having a distal bifurcation through the introducer sheath, constraining the distal bifurcation in a low-profile configuration, advancing the constrained distal bifurcation from the introducer sheath toward the renal arteries while the bifurcation remains constrained, and releasing the distal bifurcation from the constrained
15 low-profile configuration to allow entry of a first distal extension of the distal bifurcation into one of the renal arteries and a second distal extension of the distal bifurcation into the other renal artery. In some aspects, constraining the distal bifurcation in the low-profile configuration includes constraining the distal bifurcation with a sheath, a ring capture system, or a guide wire ring. In some aspects, passing the constrained distal bifurcation from the
20 introducer sheath toward the renal arteries includes advancing the sheath over a guide wire, advancing the ring capture system along a guide wire, or advancing a guide wire ring along a guide wire. In a related aspect, the method can also include advancing a second catheter through the introducer sheath, and performing an intervention with the second catheter.

[0029] In another aspect, the present invention provides a system for delivering treatment
25 to the renal arteries. The system can include a delivery catheter having a distal bifurcation, and an introducer assembly having an introducer sheath in operative association with a Y-hub. The introducer sheath can have a length in the range from about 5 cm to about 25 cm. The Y-hub can have a first port for receiving the delivery catheter and a second port for receiving a second catheter. The system can also include a constraint assembly for holding
30 the distal bifurcation of the delivery catheter in a low-profile configuration when it is advanced distally beyond the introducer sheath. In some aspects, the constraint assembly may include a jacket sheath for holding the distal bifurcation of the delivery catheter in a low-profile configuration when it is advanced distally beyond the introducer sheath. The jacket

sheath can include a guide for receiving a guide wire. The jacket sheath can also include a split tube. In some cases, the jacket sheath includes distal and proximal flow apertures. In some cases, the jacket sheath can be in sealed cooperation with the delivery catheter.

Relatedly, the delivery catheter can include a delivery catheter port for delivering an infusate to an interior of the jacket sheath. In some cases, the constraint assembly can include a collapsible ring capture system or a guide wire ring.

[0030] In another aspect, the present invention provides a method of positioning a delivery catheter in a branch lumen extending from a main lumen in a body of a patient. The method can include positioning an introducer sheath in the main lumen, advancing a delivery catheter having a distal bifurcation through the introducer sheath, constraining the distal bifurcation in a low-profile configuration, advancing the constrained distal bifurcation from the introducer sheath toward the branch lumen, and releasing the distal bifurcation from the constrained low-profile configuration to allow entry of a first distal extension of the distal bifurcation into the branch lumen. In some cases, advancing the introducer sheath in the main lumen can include advancing the introducer sheath through a puncture in a first femoral artery of the patient. The method can also include advancing a guide catheter through the introducer sheath toward a second femoral artery of the patient via the aortic bifurcation. In some cases, constraining the distal bifurcation in the low-profile configuration can include constraining the distal bifurcation with a sheath, a ring capture system, or a guide wire ring.

[0031] For a fuller understanding of the nature and advantages of the present invention, reference should be had to the ensuing detailed description taken in conjunction with the accompanying drawings.

BRIEF DESCRIPTION OF THE DRAWINGS

[0032] Fig. 1 illustrates a renal delivery system according to one embodiment of the present invention.

[0033] Fig. 2 illustrates a renal delivery system according to one embodiment of the present invention.

[0034] Fig. 3A illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0035] Fig. 3B illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0036] Fig. 4 illustrates a renal delivery system according to one embodiment of the present invention.

5 [0037] Fig. 5A illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0038] Fig. 5B illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

10 [0039] Fig. 6A illustrates a jacket sheath according to one embodiment of the present invention.

[0040] Fig. 6B illustrates a jacket sheath according to one embodiment of the present invention.

[0041] Fig. 6C illustrates a cut away view of a jacket sheath according to one embodiment of the present invention.

15 [0042] Fig. 7A illustrates a constraint assembly according to one embodiment of the present invention.

[0043] Fig. 7B illustrates a constraint assembly according to one embodiment of the present invention.

20 [0044] Fig. 7C illustrates a constraint assembly according to one embodiment of the present invention.

[0045] Fig. 8 illustrates an introducer assembly according to one embodiment of the present invention.

[0046] Fig. 9A illustrates a delivery catheter according to one embodiment of the present invention.

25 [0047] Fig. 9B illustrates a delivery catheter according to one embodiment of the present invention.

[0048] Fig. 10A illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0049] Fig. 10B illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0050] Fig. 11 illustrates a constraint assembly according to one embodiment of the present invention.

5 [0051] Fig. 12A illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0052] Fig. 12B illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

10 [0053] Fig. 12C illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0054] Fig. 13 illustrates a constraint assembly according to one embodiment of the present invention.

[0055] Fig. 14A illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

15 [0056] Fig. 14B illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0057] Fig. 15A illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

20 [0058] Fig. 15B illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0059] Fig. 16A illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0060] Fig. 16B illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

25 [0061] Fig. 17A illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0062] Fig. 17B illustrates a delivery catheter and constraint assembly according to one embodiment of the present invention.

[0063] Fig. 18 illustrates an introducer assembly according to one embodiment of the present invention.

DETAILED DESCRIPTION OF THE INVENTION

5 [0064] As discussed herein the present invention can be provided to the physician as an integrated catheter-sheath system, and can eliminate the operational step of determining the proper sheath sizing to use for a particular patient. Alternatively, the various components of the system may be provided independently or in different combinations with each other or with other conventional catheter system components. The present invention is well suited for
10 both coronary and contralateral procedures.

[0065] The description herein provided relates to medical material delivery systems and methods in the context of their relationship in use within a patient's anatomy. Accordingly, for the purpose of providing a clear understanding, the term proximal should be understood to mean locations on a system or device relatively closer to the operator during use, and the
15 term distal should be understood to mean locations relatively further away from the operator during use of a system or device. These present embodiments below therefore generally relate to local renal drug delivery generally from the aorta; however, it is contemplated that these systems and methods may be suitably modified for use in other anatomical regions and for other medical conditions without departing from the broad scope of various of the aspects
20 illustrated by the embodiments. Applicants have now completed over one hundred human renal clinical cases with the delivery system of the present invention.

[0066] In general, the disclosed material delivery systems will include a fluid delivery assembly, a proximal coupler assembly and one or more elongated bodies, such as tubes or catheters. These elongated bodies may contain one or more lumens and generally consist of a
25 proximal region, a mid-distal region, and a distal tip region. The distal tip region will typically have means for delivering a material such as a fluid agent. It is appreciated, however, that the present systems may be configured to deliver any of a wide variety of treatment modalities, including the therapeutic application of ultrasound and other types of treatment energy. Radiopaque markers or other devices may be coupled to the specific
30 regions of the elongated body to assist introduction and positioning.

[0067] The material delivery system is intended to be placed into position by a physician, typically either an interventionalist (e.g. a cardiologist or radiologist) or an intensivist, a

physician who specializes in the treatment of intensive-care patients. The physician will gain access to a femoral artery in the patient's groin, typically using a Seldinger technique of percutaneous vessel access or other conventional method.

[0068] For additional understanding, further more detailed examples of other systems and methods for providing local renal drug delivery are variously disclosed in the following published references: WO 00/41612 to Keren et al.; and WO 01/83016 to Keren et al. The disclosures of these references are herein incorporated in their entirety by reference thereto. Moreover, various combinations with, or modifications according to, various aspects of the present embodiments as would be apparent to one of ordinary skill upon review of this disclosure together with these references are also considered within the scope of invention as described by the various independently beneficial embodiments described below.

[0069] Turning now to the drawings, **Fig. 1** illustrates a renal delivery system 10 according to one embodiment of the present invention. System 10 includes an introducer assembly 100, a renal delivery catheter 200, and a coronary guide catheter 300 suitably for introducing, for example, a balloon angioplasty catheter (not shown). Introducer assembly 100 can have a Y-hub 110 and an introducer sheath 120. Typically, Y-hub distal end 112 is coupled with introducer sheath proximal end 124. Delivery catheter 200 includes a delivery catheter shaft 201 having a distal end 202 and a proximal end 204. Renal delivery catheter shaft 201 is partially disposed within introducer assembly 100, such that delivery catheter shaft distal end 202 extends from introducer sheath distal end 122, and delivery catheter shaft proximal end 204 extends from a first port 116 of Y-hub proximal end 114. Similarly, coronary guide catheter 300 is partially disposed within introducer assembly 100, such that coronary guide catheter distal end 302 extends from introducer sheath distal end 122, and coronary guide catheter proximal end 304 extends from a second port 118 of Y-hub proximal end 114. Introducer sheath 120 can have a length selected to terminate well below the renal arteries, preferably below the aortic branch 50. The length is typically in the range from about 5 cm to about 25 cm, typically from about 10 cm to about 15 cm. In many cases, introducer sheath 120 will have a very thin wall. The wall may be as thin as possible or practical, which can allow for an optimal vessel entry profile and a greater inner diameter. In some cases, the outer diameter of introducer sheath 120 may be, for example, less than or equal to 1 French. In other cases, the outer diameter may be about 2 French. In related cases, the outer diameter of introducer sheath may be within the range of between about 0.005" to about .0025", or within the range of between about 0.009" to about 0.015".

[0070] Introducer assembly 100 allows for the placement of renal delivery catheter 200 and coronary guide catheter 300 (as well as associated interventional catheters) into a patient's vasculature via a single vessel entry puncture 90. In some cases, system 10 can be used for renal drug infusion during a primary catheterization procedure, such as a coronary intervention. Advantageously, introducer sheath 120 can simultaneously facilitate introduction of infusion catheter 200 for a renal infusion procedure, and introduction of coronary guide catheter 300 for a coronary procedure. Typically, introducer sheath 120 is advanced through single puncture 90 in a femoral artery 30 or an iliac artery 40 of the patient, below or caudal to the aortic bifurcation 50. Delivery catheter 200 and coronary guide catheter 300 are then inserted through introducer assembly 100, into the patient's body, to their respective desired locations. **Fig. 1** illustrates delivery catheter 200 having a distal bifurcation 210 in a deployed high-profile configuration. As further discussed below, system 10 can include a jacket sheath 610 for constraining a distal bifurcation 210 of delivery catheter 200 in a low-profile position.

[0071] The present inventors have discovered that an introducer sheath 120 having the above-described dimensions can provide the physician with many practical operational advantages. For example, the physician can leave introducer sheath 120 in place while retracting and advancing jacket sheath 610, thereby simplifying the cannulation procedure. Moreover, a short introducer sheath 120 allows for a one-size-fits-all approach for placing delivery catheter 200 in the patient's body, as the system does not require an introducer sheath that is optimally sized in each case according to the patient's anatomy (e.g. the distance from the puncture site to the renal artery origins or ostia). Such features are highly amenable to improved standardization in case planning. Universal sizing can eliminate the need to use introducer sheaths that are pre-sized to fit the patient's anatomy.

[0072] Relatedly, the present invention avoids the need to have excess external sheath length extending proximally from the entry site, as may be the case where the distance between the puncture site and the deployment site is significantly less the length of standard introducer sheaths. And where it is necessary to retract introducer sheath 120 from the patient, the reduced length extending proximally from the patient will create less interference where the physician may need to maneuver other components of device 10. By reducing the length of introducer sheath 120 that extends proximally from the patient, it is possible to avoid or minimize the risk of having an inadequate length for an auxiliary catheter such as a coronary catheter to reach its intended site. In other words, a shortened introducer sheath 120

can allow Y-hub 110 to be disposed closer to a vessel entry point such as puncture 90. In general, coronary guide catheters are sized to span the typical distance from a vessel entry site to the coronary arteries. Using a shortened introducer sheath 120 avoids the situation where a guide catheter is coupled with introducer assembly 100 but remains too far from the groin entry, such that during the procedure the proximal end of the guide catheter engages Y-hub 110 before the distal end of the guide catheter reaches the target coronary vessel. Auxiliary components of device 10 will therefore encounter less interference or friction from introducer sheath 120 within the interior of the patient's vessel. Advantageously, use of the present invention eliminates many of the difficulties associated with managing iliac, aortic, and other vascular tortuosity.

[0073] Standard length introducer sheaths are often placed directly in the aorta, and catheter devices are continuously passed therethrough. Advantageously, the shortened introducer sheath dimensions of the present invention minimize the potential for clot formation within introducer sheath 120 and Y-hub 110. Thus, the present systems can eliminate or reduce the need for a constant saline or heparinized saline drip, which is intended to prevent clotting in introducer sheath 120. Relatedly, in some cases there can be an increased risk of clots dislodging when adjunctive commercial products are passed through standard length introducer sheaths, particularly at advancement. The introducer sheath 120 of the present invention minimizes such risks. The present invention confers such benefits while still providing a full-covered advancement of distal bifurcation 210 to the intended deployment site. Accordingly, the present invention provides a system with improved stability characteristics.

[0074] The length of introducer sheath 120 can be such that delivery catheter 200 and coronary catheter 300 can both exit introducer sheath 120 well below the renal arteries. Thus, there is less opportunity to displace delivery catheter 200 when advancing coronary catheter 300. Similarly, because a distal bifurcation 210 of delivery catheter 200 is typically not maintained within introducer sheath 120, it is unlikely that significant friction or other jamming complications will develop between coronary guide catheter 300 and delivery catheter 200. This is true whether delivery catheter distal end 210 is in a deployed or undeployed configuration. In other words, the length of introducer sheath 120 can allow adjunctive coronary catheter 300 to exit introducer sheath 120 at a sufficient distance from renal arteries 60, so as to provide improved stability of delivery catheter 200 as it interacts with renal arteries 60. Because the length of introducer sheath 120 reduces the need to

manage vessel tortuosity and possible interference with other system components, the need for substantial columnar and radial support for sheath 120, for example from coil reinforcements, can be eliminated or greatly reduced. Introducer sheath 120 can be prepared with a thin wall extrusion, where the thickness of the wall is less than currently used multi-layer and reinforced sheath tubes.

[0075] In addition to facilitating various coronary procedures, the present invention can also be used in conjunction with a wide variety of peripheral procedures. In one example, the present invention can be used in contralateral superficial femoral artery (SFA) vessel advancement for critical limb salvage cases, which may be particularly useful in treating diabetic patients. Similarly, the present invention can be used to effect various procedures in the abdominal or femoral arteries, and can be used to treat occlusive peripheral vascular disease, critical limb ischemia, and other related conditions.

[0076] Fig. 2 illustrates a renal delivery system 10 according to one embodiment of the present invention which is well suited for such peripheral procedures. Toward this end, system 10 can include an introducer assembly 100, a delivery catheter 200, and a peripheral guide catheter 400. Peripheral guide catheter 400 is partially disposed within introducer assembly 100, such that peripheral guide catheter distal end 402 extends from introducer sheath distal end 122, and peripheral guide catheter proximal end 404 extends from a second port 118 of Y-hub proximal end 114. A renal infusion may be performed simultaneously with a contralateral iliac vessel procedure via a single vessel entry 90 through introducer assembly 100. Because introducer sheath distal end 122 can sit below the aortic bifurcation 50, system 10 can be used in "up-and-over" procedures where contralateral access is desired. Consequently, there is no need to retract introducer sheath 120 to situate its distal end below the aortic bifurcation 50, and similarly, there is no need to perform additional manipulations of peripheral catheter 400 at Y-hub 110 which could otherwise be far from the vessel entry site if a standard length introducer sheath were used. Introducer sheath 120 provides optimal use of the working length of peripheral catheter 400, which will often be helpful for extremely distal procedures, and provides optimal handling characteristics to peripheral catheter 400. Thus, the present invention provides a system that can be universally used for femoral-based coronary procedures and peripheral procedures alike. Present system 10 can reduce friction on peripheral catheter 400, as the length of introducer sheath 120, through which both delivery catheter 200 and peripheral catheter 400 pass, is minimized.

[0077] In some embodiments, delivery catheter 200 can be used to deliver a therapeutic or diagnostic infusate to the renal arteries. It is also appreciated that delivery catheter 200 can be used to effect a wide variety of other therapeutic and diagnostic modalities at or near the renal ostia 57 or renal arteries 60, including stent placement, therapeutic energy delivery, and the like.

[0078] The present invention also provides means for maintaining distal bifurcation 210 of delivery catheter 200 in a constrained or otherwise undeployed or protected configuration.

Figs. 3A and 3B illustrate a constraint assembly 600 according to one embodiment of the present invention. Constraint assembly 600 includes a jacket sheath 610 coupled with a lead 630, which may in turn be coupled with a connector 620. **Fig. 3A** depicts delivery catheter 200 in an unconstrained or deployed configuration. Jacket sheath 610 is retracted from delivery catheter distal bifurcation 210. Delivery catheter 200 may be introduced via a sacrificial long thin-wall covering jacket 610, which can be removed after deployment. In such cases, delivery catheter 200 can be retracted without being constrained by jacket sheath 610. As shown in **Fig. 3B**, by advancing lead 630 in a distal direction along delivery catheter shaft 201, jacket sheath 610 envelops and forces distal bifurcation 210 into a collapsed, undeployed, low profile configuration. In this way, lead 630 can be retracted or advanced along the length of delivery catheter shaft 201 so as to deploy or undeploy distal bifurcation 210. In some cases, distal jacket sheath 610 may also be referred to as a movable cover jacket or capture sheath. Often, the diameter of jacket sheath 610 will be only as large as necessary to contain distal extensions or tips 212 of delivery catheter 200.

[0079] Typically, delivery catheter distal end 210 is in the undeployed configuration when it is inserted into the patient's body via introducer assembly 100. With supplemental reference to **Fig. 1**, for example, the physician may advance the constrained delivery catheter distal end 210 through the patient's femoral artery 30, through the iliac artery 40, past the aortic branch 50, and through the aorta 55 to the desired deployment location at or near the renal ostia 57. The physician can then manipulate constraint assembly 600 so as to deploy distal bifurcation 210, thereby initiating a renal cannulation procedure. Often, distal bifurcation 210 includes distal extensions 212, and deployment involves engaging distal extensions 212 with renal arteries 60. By advancing an undeployed delivery catheter distal end 210 into the patient's body, it is possible to avoid causing vessel trauma with distal extensions 212. It is believed that in many cases, it may be difficult to advance delivery catheter distal end 210 in the deployed configuration. The collapsed and low-profile state

facilitates advancement of the distal end distal extensions 212 through the patient's vasculature.

[0080] Constraint assembly 600 may be designed and used in a number of ways. For example, as seen in **Fig. 3A**, lead 630 can include a thin-wall outer membrane that sheathes delivery catheter shaft 201. In this embodiment, lead 630 is intended to remain intact, as a movable outer member over shaft 201 in a coaxial fashion. By advancing or retracting outer member lead 630, cover jacket 610 captures or releases distal extensions 212 of distal bifurcation 210. In this sense, the deployed configuration refers to the open or high-profile configuration. Incorporation of outer member lead 630 can add functional outer diameter to delivery catheter shaft 201, and may in some cases require a smaller catheter shaft 201 if the outer diameter of lead 630 is equal to or smaller than the original diameter of shaft 201. Relatedly, incorporation of outer member 630 may require an upsizing in the functional diameter of the catheter shaft 201.

[0081] It is appreciated that the present invention contemplates a wide variety of means for reversibly covering or restraining distal extensions 212 in their collapsed state. As further discussed below, constraint assemblies can include various combinations of jackets, sheaths, sleeves, covers, control wires or rods, and the like. It is further appreciated the present invention may include or otherwise be in operative association with guidance assemblies such as guide wires so as to ensure desired placement of device 10 in the patient's body. Guide wire guidance can allow guided advancement of delivery catheter 200 and/or guide catheter 300, 400 during deployment, cannulation, and removal. In some cases, guide wire guidance may not be required for device removal. For example, device removal may simply involve retracting distal extensions 212 into jacket 610, and atraumatically withdrawing collapsed bifurcated end 210 through the vasculature and out of introducer sheath 120 and entry site.

[0082] Device 10 may be retracted in either the collapsed or uncollapsed configuration. In some cases, distal extensions 212 are sufficiently soft and pliable to be retracted in an uncollapsed configuration from the patient's renal ostia 57 without causing damage, as shown in **Fig. 4**. For example, in some percutaneous applications, it is often sufficient to remove delivery catheter 200 while distal extensions 212 are still in the uncollapsed configuration. In other cases, a physician will remove device 10 in the collapsed atraumatic configuration to reduce the potential of vessel trauma and displacement of plaques, thrombi, and the like.

[0083] Figs. 5A and 5B illustrate a particular embodiment of the present invention that uses a monorail guidance system. Delivery catheter 200 shown in Fig. 5A is undeployed, as jacket sheath 610 of constraint assembly 600 maintains distal bifurcation 210 in a low profile configuration. Jacket sheath 610 includes a guide 612 adapted to slidably receive a guide wire 700. Here, guide 612 includes a longitudinal tube affixed with the body of sheath 610, in such a way that guide wire 700 is disposed laterally to the internal lumen of sheath 610. It is appreciated that in other embodiments, guide 612 may include other means for slidably engaging guide wire 700. When delivery catheter 200 is advanced through the patient's artery, guide 612 slides along guide wire 700, while at the same time remaining fixed relative to delivery catheter distal bifurcation 210. In one embodiment, lead 630 includes a control wire or rod instead of an axially disposed cylindrical outer member, and thus is not of a full coaxial design over the delivery catheter shaft 201. Cover jacket 610 may be coaxially located on delivery catheter shaft 201, which remains permanently over delivery catheter shaft 201. Delivery catheter 200 shown in Fig. 5B is in a deployed configuration, as jacket sheath 610 is retracted away from distal bifurcation 210. Typically, this retraction also involves sliding guide 612 in a proximal direction along guide wire 700, although in some embodiments retraction of constraint assembly 600 will be accompanied by retraction of guide wire 700. In some cases, guide 612 may include a series of segmented tubular segments. In related cases, guide 612 may include a partially tubular segment having a C-shaped cross section, whereby guide 612 may be snapped onto and off of guide wire 700.

[0084] As discussed above, jacket sheath 610 may be coupled with lead 630 for advancement and retraction. It is appreciated that lead 630 can include any of a variety of control wires, or small diameter shafts or rods. In some iterations, lead 630 is permanently attached with jacket sheath 610, and in others it is removably coupled with jacket sheath 610. In some cases, jacket sheath 610 can be positioned just outside of introducer assembly Y-hub 110 without an attached lead 630. This configuration is facilitated by the option of the ability to remove or detach lead 630 from jacket sheath 610. On the other hand, the option of having lead 630 permanently connected with cover jacket sheath 610 can provide simplicity in manufacture as well as potentially high strength and ease of use. In some embodiments, lead 630 is approximately 35 cm in length, and sufficiently narrow in diameter so as not to interfere with the catheterization procedure. Another design for the detachable lead 630 uses a means of connection of the cover jacket to currently available wires such as a guide wire, and one is meant for a dedicated control wire, with specific means of connection to the jacket.

[0085] As shown in **Fig. 6A**, proximal end 611 of jacket sheath 610 may include an aperture 612a for receiving guide wire (not shown), such that a segment of guide wire is housed within the sheath 610. Here, guide 612 resides substantially within the cylindrical wall of jacket sheath 610. In the embodiment shown in **Figs. 6B and 6C**, jacket sheath 610 includes two lumens 616 for housing distal extensions 212, and a lumen 618 for housing guide wire 700.

[0086] **Figs. 7A and B** illustrate another embodiment of constraint assembly 600 which includes a split tube type of jacket sheath 610. Guide wire 700 may be passed through split 619, instead of being threaded through a monorail-type guide. Similarly, delivery catheter shaft 201 or guide catheters 300, 400 may be passed through split 619, thus allowing for easy removal and replacement of jacket sheath 610. As depicted in **Fig. 7C**, in some cases jacket sheath 610 may not be continuous, but instead may include multiple segments. Such a configuration can be advantageous by providing less material for introduction into the vasculature, thus diminishing the risk for clotting. Relatedly, by providing less luminal surface, the amount of friction between jacket sheath 610 and delivery catheter 200 or adjunctive catheter 300, 400.

[0087] As seen in **Fig. 8**, jacket sheath 610 can be passed through first port 116 of Y-hub 110. Lead 630 can be manipulated by a physician to control movement of jacket sheath 610. By allowing jacket 610 to be fully retracted out of introducer sheath 120 and Y-hub 110, external to the patient's vessel entry 90, the functional diameter of device 10 remaining in the patient (e.g. catheters 200, 300, or 400) can be reduced. Often, jacket sheath can remain outside of the patient's body during the dwell period. Relatedly, larger diameter delivery catheters 200, adjunctive coronary catheters 300, or peripheral catheters 400 may be used in devices 10 having such fully retractable jackets 610. For the same reason, introducer sheaths 120 having smaller outside diameter may be used. Jacket sheath 610 may include markers 617 for position verification. In some embodiments, markers 617 may include radiopaque materials.

[0088] **Figs. 9A and 9B** illustrate a variable shape distal arm design for delivery catheter 200. Catheter 200 may include a bifurcated pull wire 205 that can be affixed with the inside distal tips of distal extensions 212, on the side of each distal extension 212 that becomes the inside of the curve. Thus, when tension is put on pull wire 205, it effectively shortens the side of distal extension 212 it is attached to, causing it to bend. This deployed orientation,

where distal extensions 212 are bent and separated from one another, is shown in **Fig. 9A**. Releasing pull wire 205 allows distal extensions 212 to go back to their default at-rest straight configuration. This collapsed orientation is shown in **Fig. 9B**.

[0089] **Figs. 10A and 10B** illustrate another embodiment of the present invention, where delivery catheter 200 includes a tapered, atraumatic distal tip or cone 750, or other dilator means for maximizing the functional cross-section of a vessel prior to deployment delivery catheter 200. In some cases, distal tip 750 is not coupled with delivery catheter 200. In some cases, distal tip 750 may have a guide 752 such as a rail or a lumen which can be used in conjunction with guide wire 700. It will be appreciated that guide wire 700 can be a floppy-type wire, which can reduce the likelihood of tapered cone 750 creating a vessel perforation when advanced for distal catheter arm cannulation. The diameter of the guide wire may vary, although in many cases is it about 0.035 inches. **Fig. 10A** shows distal extensions 212 of delivery catheter 200 in a high-profile deployed configuration. Jacket sheath 610 may include a seal or gasket 611 which is mounted on or otherwise connected with catheter shaft 201, and therefore advances and retracts with delivery catheter shaft 201 advancement and retraction while remaining in sealed engagement with inner surface of jacket sheath 610. Here, gasket 611 is disposed toward the distal end of jacket sheath 610, and the proximal end of jacket sheath is otherwise sealed. Thus, jacket sheath 610 can be closed to blood flow when distal bifurcation 210 is in the deployed condition. **Fig. 10B** shows distal extensions 212 of delivery catheter 200 in a low-profile undeployed configuration. Here, gasket 611 is disposed toward the proximal end of jacket sheath 610, and distal cone 750 effectively seals the distal end of jacket sheath 610. By preventing blood from flowing through jacket sheath 610, gasket 610 can help reduce the risk of thrombus formation within sheath 610.

[0090] Cover jacket sheath 610 for constraining distal distal extensions 212 may be of multiple configurations, depending on the overall product needs. For example, jacket sheath 610 can accommodate a guide wire lumen or monorail-type arrangement. Further, jacket sheath 610 may be designed with tapers on each end or, as shown in **Fig. 11**, may be include an untapered proximal end and a tapered distal end. Here, a distal end of guide 612 exits at a central tip of jacket sheath 610, whereas distal bifurcation 210 exits lateral to the central tip. Such configurations can provide less traumatic passage of delivery catheter 200 through the patient's vasculature.

[0091] The present invention provides a variety of approaches for preventing or inhibiting the formation of a blood clot or thrombus within jacket sheath 610, for example by avoiding the pooling of stagnant or static blood therein. In some cases, these approaches will include a jacket sheath 610 that remains integral with delivery catheter shaft 201. As shown in **Fig.**

5 **12A**, jacket sheath 610 can be in sealed cooperation with delivery catheter shaft 201 so as to prevent blood from flowing into or through sheath 610. Gasket 611 can prevent blood infusion, or infusate seepage, at the distal end of jacket sheath 610, and the proximal end of sheath 610 may be otherwise sealed with shaft 201. Gasket 611 can be fixed with shaft 201, and in sealed engagement with the inner surface of jacket sheath 610. **Fig. 12B** illustrates
10 delivery catheter shaft 200 having a port 206 in fluid communication with interior of jacket sheath 610. Port 206 can be used to deliver infusate from delivery catheter shaft 201 to sheath 610. By allowing infusate to fill the luminal space or effective cavity of jacket sheath 610, a positive infusate flow into this cavity can be created, so as to prevent or inhibit thrombus formation. Similarly, the interior chamber of jacket sheath 610 can be pressurized
15 with the infusate from port 206. Often, jacket sheath 610 will be sealed with one or more gaskets or seals 611 so as to prevent blood infusion or infusate seepage. A positive pressure of an effective reservoir of infusate can build up in the cavity remaining in jacket sheath 610. Because there is a slight pressure (greater than blood pressure), the blood cannot so easily displace the infusate in the cavity, and thus it prevents clotting. In a nonsealed or open
20 version as seen in **Fig. 12C**, jacket sheath 610 may include proximal end vents or perforations 613 so as to allow continuous blood flow through jacket sheath 610. Here, incoming blood perfuses out of sheath 610 via fenestrations, slots, or other openings 613 in jacket sheath 610.

[0092] **Fig. 13** illustrates a partial view of delivery catheter shaft 201 according to one
25 embodiment of the present invention. Lead 630 is shaped in a semi-circle or modified “D” shape, and nests with delivery catheter shaft 201, thus providing facile passage through introducer sheath (not shown). It is appreciated that lead 630 can have any of a variety of shapes, and many times will be configured so as to minimize the geometry or the overall functional diameter (or functional cross-sectional area) of the lead 630 and delivery catheter
30 shaft 201 combination. In a related embodiment, lead 630 can be a semi-ridged tube made of polymer or metal that travels along the length of the delivery catheter shaft 201 as a rail type system. In this sense, a modified “D” shape can be useful where there is limited need for

torque to be applied to lead 630, but lead 630 still needs to withstand compression and tension forces associated with advancement and retraction of jacket sheath 610.

[0093] **Figs. 14A and 14B** illustrates a constraint assembly 600 and delivery catheter 200 according to one embodiment of the present invention. **Fig. 14A** shows constraint assembly 600 is a retracted position, such that distal extensions 212 of delivery catheter 200 are in a deployed orientation. **Fig. 14B** shows constraint assembly 600 in an advanced position, such that distal extensions 212 of delivery catheter 200 are in an undeployed orientation. Constraint assembly 600 can be fixed in the advanced position by locking or otherwise removably coupling connector 620 with a clamp 207 of delivery catheter 200. In some cases, all components of constraint assembly 600 (e.g. handle 620, lead 630, and sheath jacket 610) can be constructed so as to allow easy advancement through Y-hub 110 (not shown here), and along guide wire 700 (not shown here).

[0094] **Figs. 15A and 15B** illustrate a constraint assembly 600 and delivery catheter 200 according to one embodiment of the present invention. Constraint assembly 600 includes a collapsible ring capture system 640 that includes a plurality of rings 640a configured to slidably engage delivery catheter 200 in a coaxial fashion. Connector 630 may be coupled with one or more rings 640a of collapsible ring capture system 640. **Fig. 15A** shows constraint assembly 600 in an advanced position, such that distal extensions 212 of delivery catheter 200 are in an undeployed configuration. **Fig. 15B** shows constraint assembly 600 in a retracted position, such that distal extensions 212 of delivery catheter 200 are in a deployed configuration. When connector 630 is retracted, rings 640a slide onto catheter hypotube or shaft 201, thus allowing distal extensions 212 to snap into their open configuration. Rings 640a can be designed such that once they are retracted onto hypotube or shaft 201, they collapse onto the surface of hypotube 201 to optimize the functional cross sectional area of the vessel.

[0095] **Figs. 16A and 16B** illustrate a means for providing constrained advancement of delivery catheter 200 to the renal ostia or other deployment locations, according to one embodiment of the present invention. **Fig. 16A** shows distal bifurcation 210 in a restrained position. Each distal extension 212 of distal bifurcation 210 includes a guide wire ring 212a coupled therewith, such that guide wire rings 212a are in slidable cooperation with guide wire 700. Guide wire rings 212a may be disposed at or near the distal tips of distal extensions 212, or at any location along the length of distal extensions 212. When distal bifurcation 210

is placed in an axially constrained state, such that distal extensions 212 are more or less aligned or parallel to one another, rings 212a can be aligned, such that guide wire 700 may be placed through rings 212a, thus holding distal extensions 212 together. In some embodiments, guide wire 700 may be a 0.035" diameter guide wire. As seen in **Fig. 16B**,
5 when delivery catheter 200 is placed in position at or about a deployment location such as the renal arteries, guide wire 700 may be withdrawn, allowing distal extensions 212 to move toward their deployed orientation. Cannulation of the renal arteries can be achieved in the manner described in previously incorporated U.S. Patent Application No. 11/084,738.

[0096] **Figs. 17A and 17B** illustrate a means for providing constrained advancement of
10 delivery catheter 200 to the peri-renal aorta or other deployment locations, according to one embodiment of the present invention. **Fig. 17A** shows distal bifurcation 210 in a restrained orientation. Constraint assembly 600 includes a spring 650, where a spring distal end 652 is coupled with jacket sheath 610, and a spring proximal end 654 is coupled with delivery catheter shaft 201. Here, spring 650 is in a relaxed state, and jacket sheath 610 remains
15 disposed distally along delivery catheter shaft 201 so as to constrain distal extensions 212 in a collapsed configuration. After insertion and deployment, spring-loaded jacket 610 can allow distal extensions 212 to be easily recaptured for withdrawal, by releasing tension in lead 630 to relax spring 650. **Fig. 17B** shows distal bifurcation 210 in a deployed configuration. When delivery catheter 200 is placed in the desired deployment location, for example in the
20 aorta, lead 630 can be actuated so as to retract jacket sheath 610 and compress spring 650, thereby allowing distal extensions 212 to open, for example, against the aortic walls.

[0097] As noted previously, embodiments that include a delivery catheter shaft or hypotube 201 in combination with wire-type lead 630 (shown in **Fig. 5A**) can often provide a smaller profile or functional cross-section diameter than embodiments that include a delivery catheter
25 shaft or hypotube 201 in combination with coaxial sleeve tube-type lead 630 (shown in **Fig. 3A**). It is appreciated that in some instances a coaxial connector sleeve tube-type lead 630 may be useful where a substantial pushing motion or compressive force is required to advance jacket sheath 610 over deployed distal extensions 212 in order to return distal bifurcation 210 to a constrained orientation. Yet the present inventors have discovered that
30 the use of a spring-loaded jacket sheath 610 as described above can remove or reduce the need to push on lead 630 to apply a compressive force to sheath 610. Relatedly, spring 650 or a similar spring-like means can provide a constant compressive load on catheter shaft 201, and a constant tensile load on lead 630. These forces can be appropriately relatively sized to

one another as desired. In some cases, delivery catheter hypotube or shaft 201 outer diameter (OD) can be in a range between about 1 French and about 3 French. In some cases, the lead 630 diameter can be in a range between about 0.010 and about 0.025", sized appropriately for delivery catheter hypotube 201.

- 5 [0098] The present invention is also well suited for use with stand-alone delivery catheter systems that do not include a guide catheter. As seen in **Fig. 18**, device 10 includes a delivery catheter 200 configured to pass through introducer assembly 100, which includes introducer sheath 120, but not Y-hub 110. This configuration may be useful in situations where a secondary adjunctive catheter 300, 400 is not needed, and thus a lower overall profile
10 can be achieved with such a stand-alone system.

[0099] While the above provides a full and complete disclosure of certain embodiments of the present invention, various modifications, alternate constructions and equivalents may be employed as desired. Therefore, the above description and illustrations should not be construed as limiting the invention, which is defined by the appended claims.

WHAT IS CLAIMED IS:

1 1. A method for positioning a delivery catheter in the renal arteries, the
2 method comprising:
3 positioning an introducer sheath in an iliac artery;
4 advancing a renal delivery catheter having a distal bifurcation through the
5 introducer sheath;
6 constraining the distal bifurcation in a low-profile configuration;
7 advancing the constrained distal bifurcation from the introducer sheath toward
8 the renal arteries while the bifurcation remains constrained; and
9 releasing the distal bifurcation from the constrained low-profile configuration
10 to allow entry of a first distal extension of the distal bifurcation into one of the renal arteries
11 and a second distal extension of the distal bifurcation into the other renal artery.

1 2. The method of claim 1, wherein constraining the distal bifurcation in
2 the low-profile configuration comprises constraining the distal bifurcation with a sheath.

1 3. The method of claim 2, wherein passing the constrained distal
2 bifurcation from the introducer sheath toward the renal arteries comprises advancing the
3 sheath over a guide wire.

1 4. The method of claim 1, wherein constraining the distal bifurcation in
2 the low-profile configuration comprises constraining the distal bifurcation with a ring capture
3 system.

1 5. The method of claim 4, wherein passing the constrained distal
2 bifurcation from the introducer sheath toward the renal arteries comprises advancing the ring
3 capture system along a guide wire.

1 6. The method of claim 1, wherein constraining the distal bifurcation in
2 the low-profile configuration comprises constraining the distal bifurcation with a guide wire
3 ring.

1 7. The method of claim 6, wherein passing the constrained distal
2 bifurcation from the introducer sheath toward the renal arteries comprises advancing the
3 guide wire ring along a guide wire.

1 8. The method of claim 1, further comprising advancing a second catheter
2 through the introducer sheath.

1 9. The method of claim 8, further comprising performing an intervention
2 with the second catheter.

1 10. A system for delivering treatment to the renal arteries, the system
2 comprising:
3 a delivery catheter having a distal bifurcation;
4 an introducer assembly comprising an introducer sheath in operative
5 association with a Y-hub, wherein the introducer sheath has a length in the range from about
6 5 cm to about 25 cm, and the Y-hub includes a first port for receiving the delivery catheter
7 and a second port for receiving a second catheter; and
8 a constraint assembly for holding the distal bifurcation of the delivery catheter
9 in a low-profile configuration when it is advanced distally beyond the introducer sheath.

1 11. The system of claim 10, wherein the constraint assembly comprises a
2 jacket sheath for holding the distal bifurcation of the delivery catheter in a low-profile
3 configuration when it is advanced distally beyond the introducer sheath.

1 12. The system of claim 11, wherein the jacket sheath comprises a guide
2 for receiving a guide wire.

1 13. The system of claim 11, wherein the jacket sheath comprises a split
2 tube.

1 14. The system of claim 11, wherein the jacket sheath comprises distal and
2 proximal flow apertures.

1 15. The system of claim 9, wherein the jacket sheath is in sealed
2 cooperation with the delivery catheter.

1 16. The system of claim 15, wherein the delivery catheter comprises a
2 delivery catheter port for delivering an infusate to an interior of the jacket sheath.

1 17. The system of claim 8, wherein the constraint assembly comprises a
2 ring capture system or a guide wire ring.

1 18. A method of positioning a delivery catheter in a branch lumen
2 extending from a main lumen in a body of a patient, the method comprising:
3 positioning an introducer sheath in the main lumen;
4 advancing a delivery catheter having a distal bifurcation through the
5 introducer sheath;
6 constraining the distal bifurcation in a low-profile configuration;
7 advancing the constrained distal bifurcation from the introducer sheath toward
8 the branch lumen; and
9 releasing the distal bifurcation from the constrained low-profile configuration
10 to allow entry of a first distal extension of the distal bifurcation into the branch lumen.

1 19. The method of claim 18, wherein advancing the introducer sheath in
2 the main lumen comprises advancing the introducer sheath through a puncture in a first
3 femoral artery of the patient, further comprising advancing a guide catheter through the
4 introducer sheath toward a second femoral artery of the patient via the aortic bifurcation.

1 20. The method of claim 18, wherein constraining the distal bifurcation in
2 the low-profile configuration comprises constraining the distal bifurcation with a member
3 selected from the group consisting of a sheath, a ring capture system, and a guide wire ring.

1/19

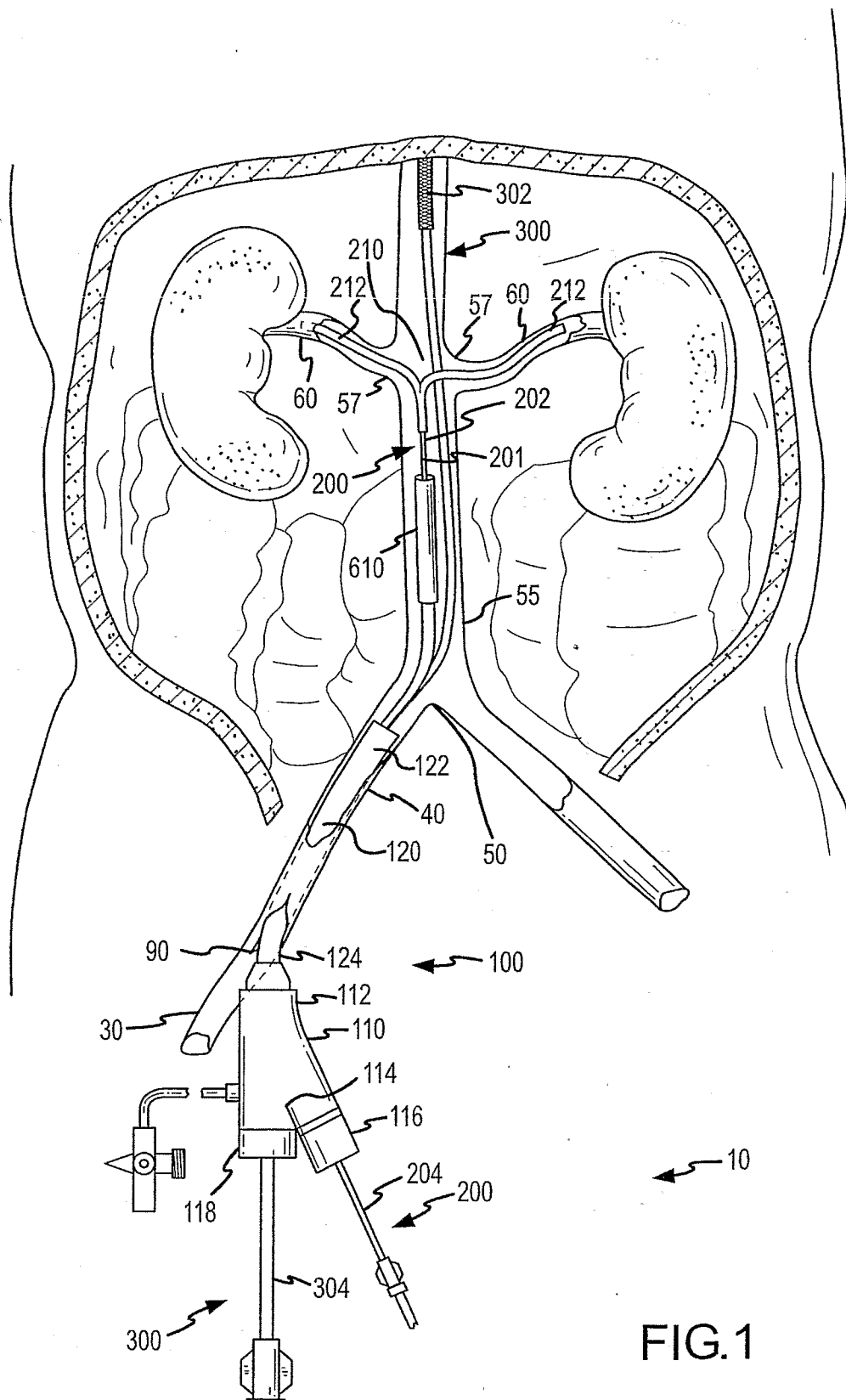


FIG.1

2/19

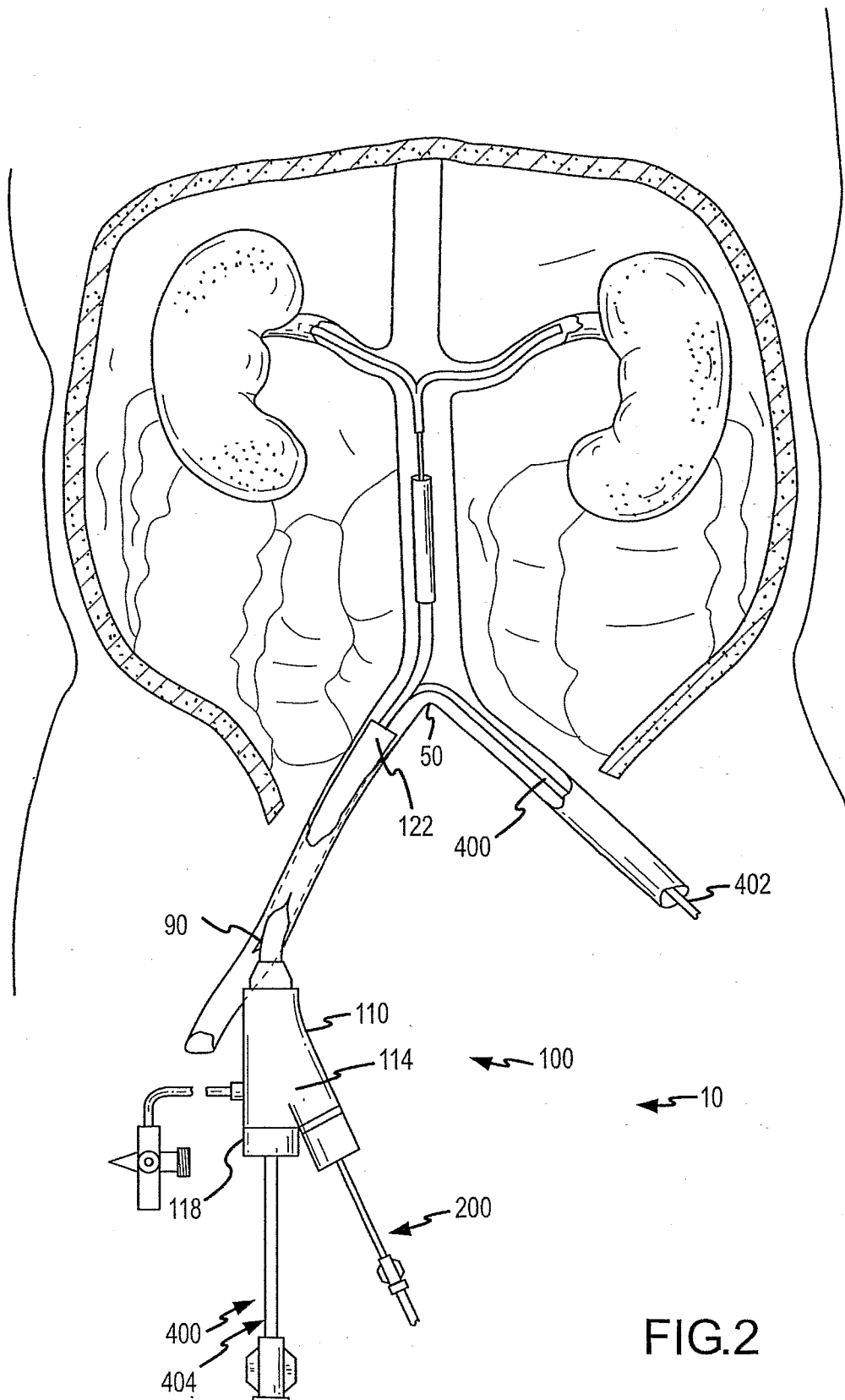


FIG.2

3/19

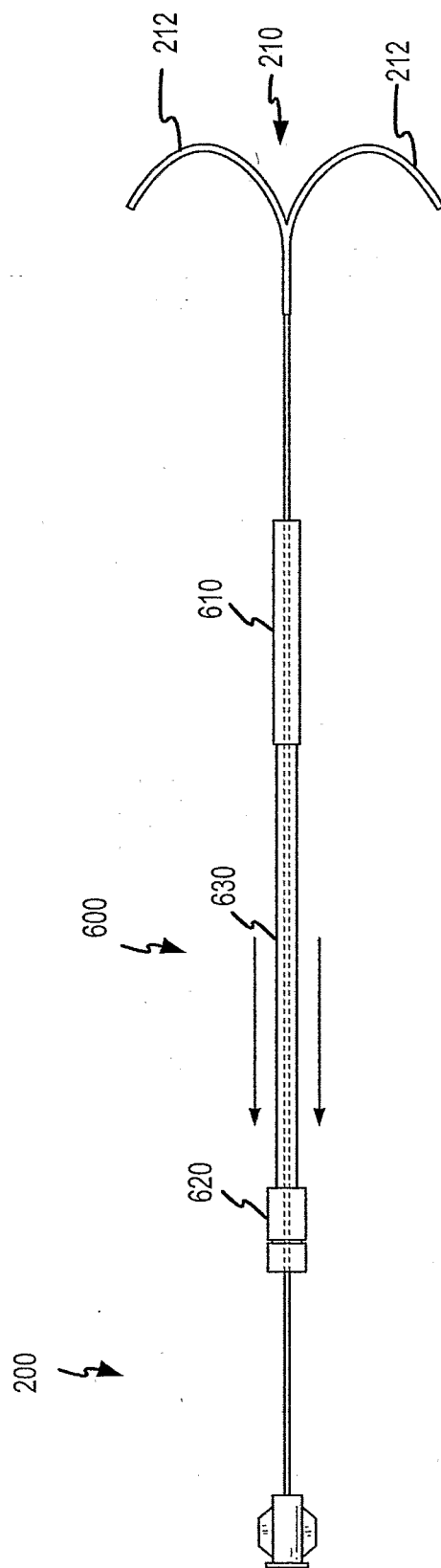


FIG. 3A

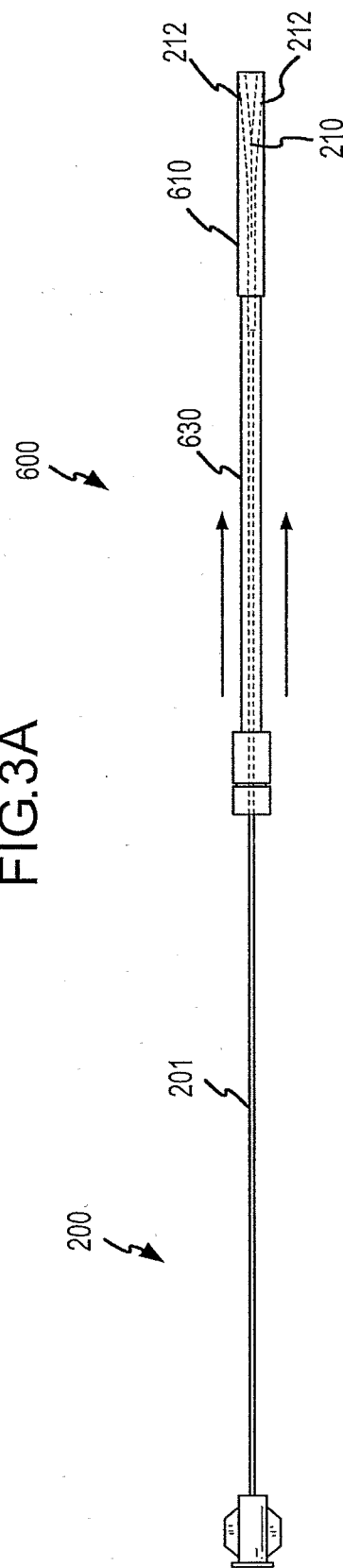


FIG. 3B

4/19

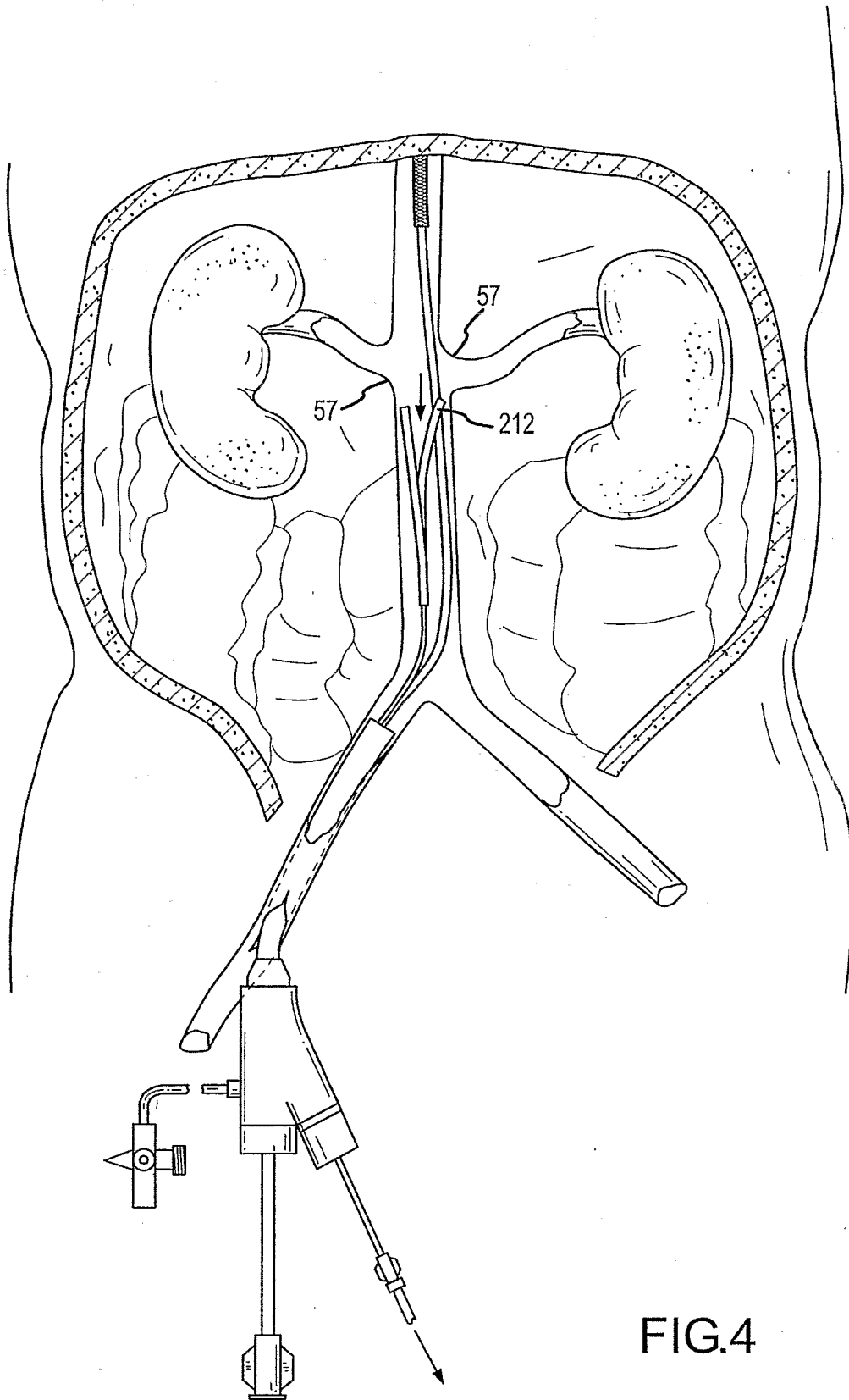


FIG.4

5/19

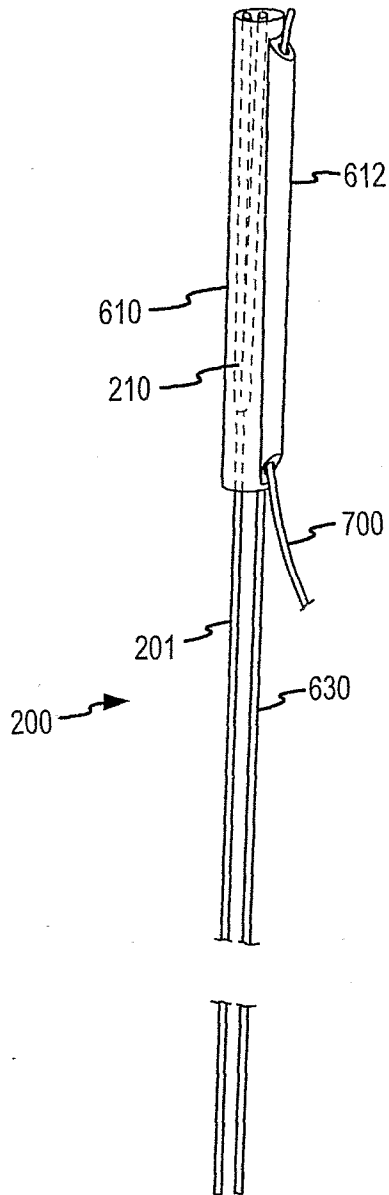


FIG. 5A

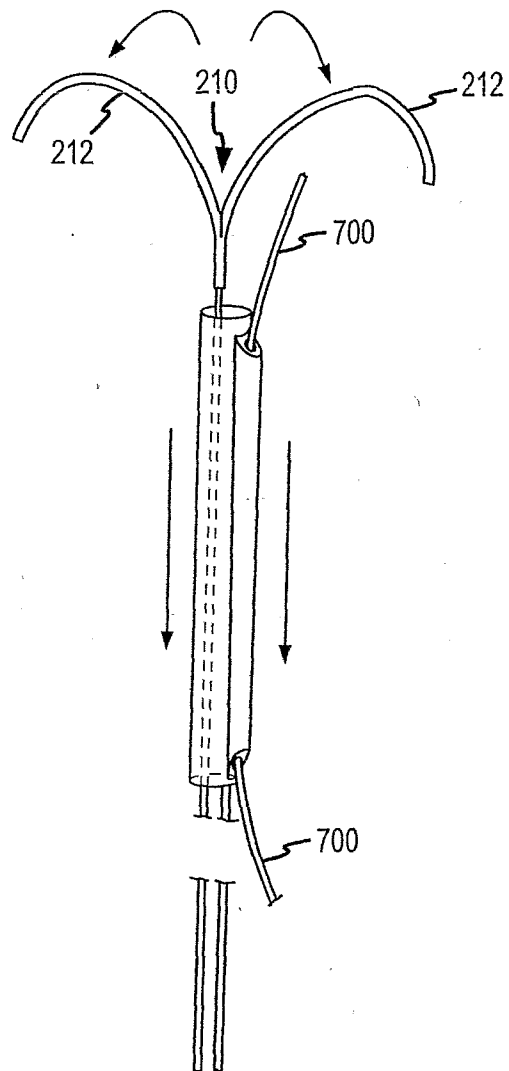


FIG. 5B

6/19

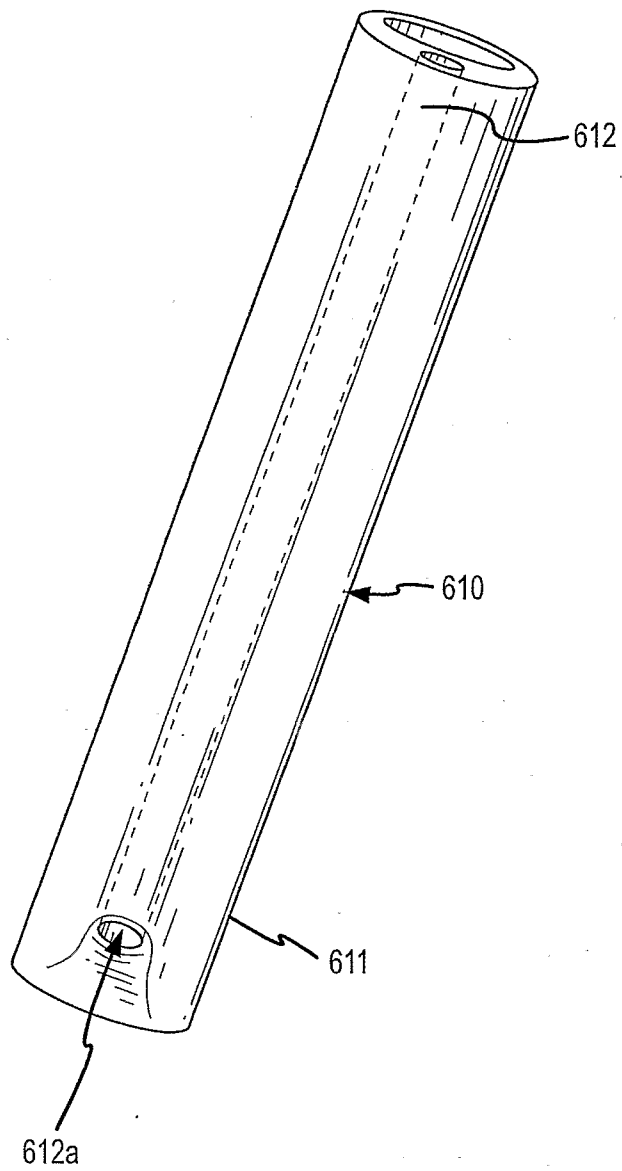


FIG. 6A

7/19

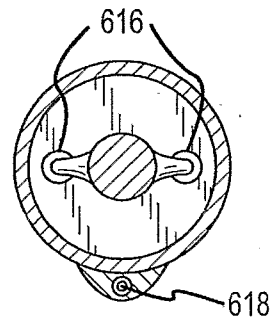
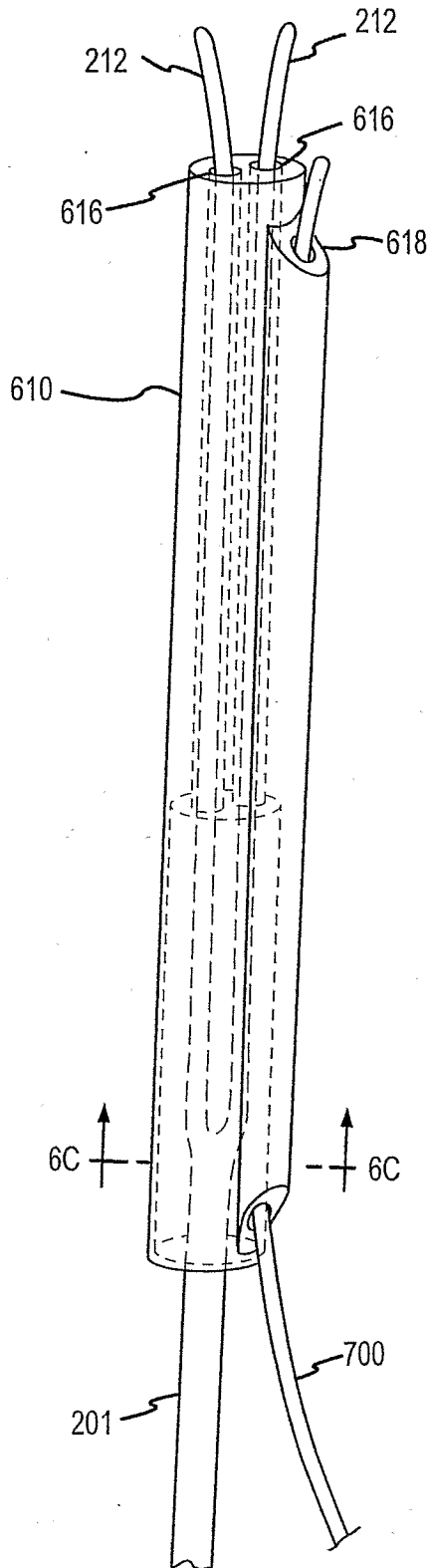


FIG. 6C

FIG. 6B

8/19

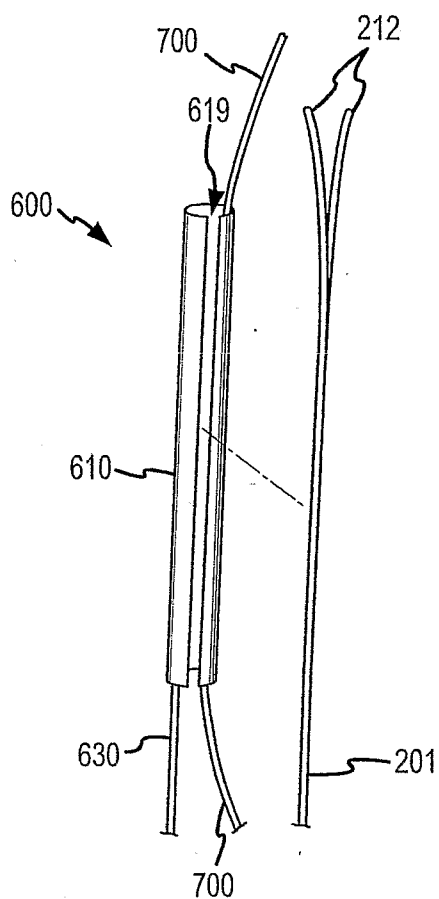


FIG. 7A

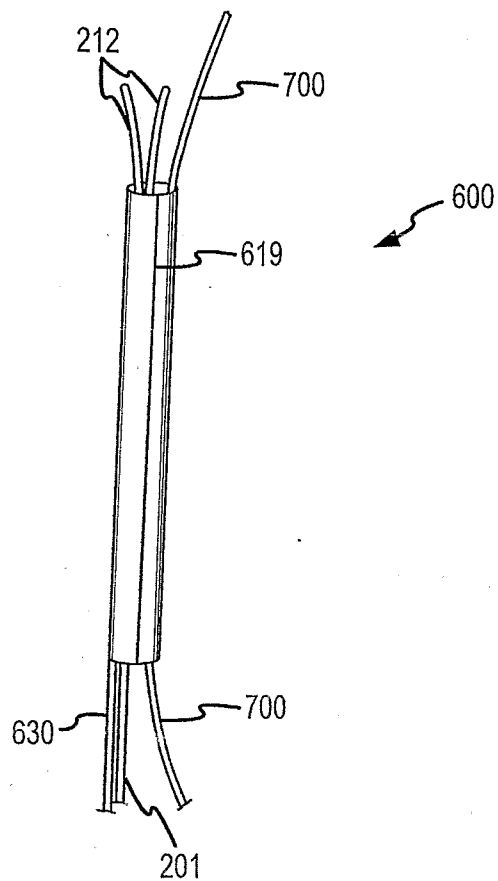


FIG. 7B

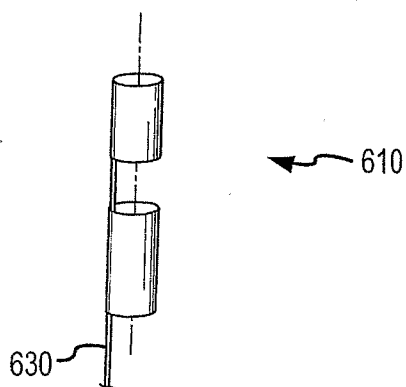


FIG. 7C

9/19

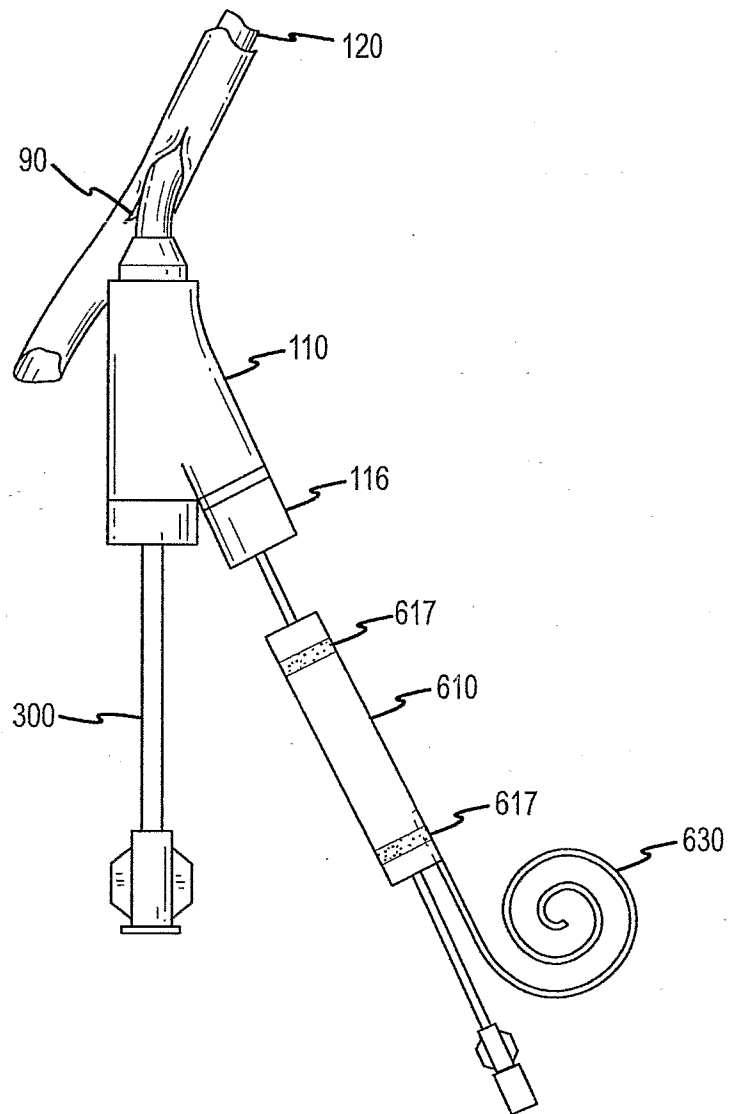


FIG. 8

10/19

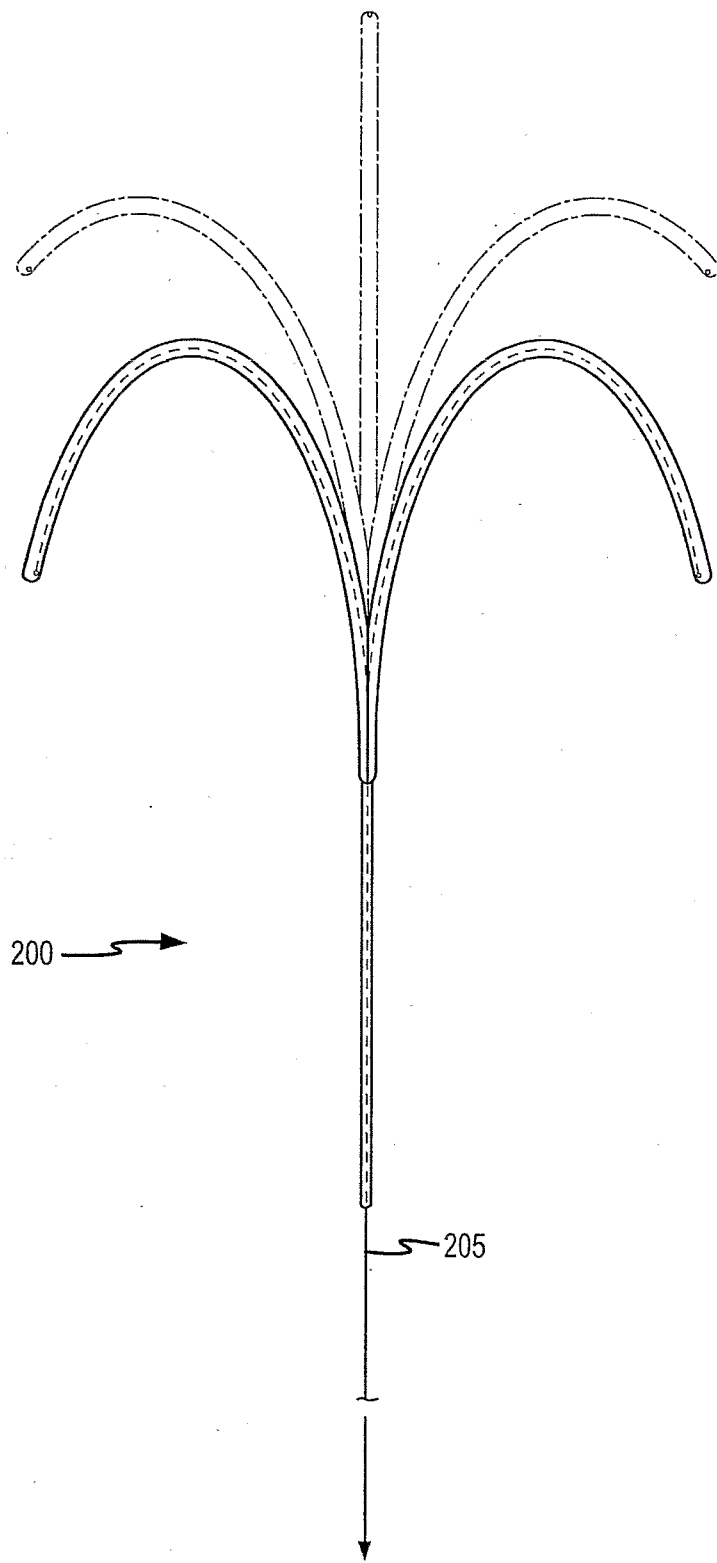


FIG. 9A

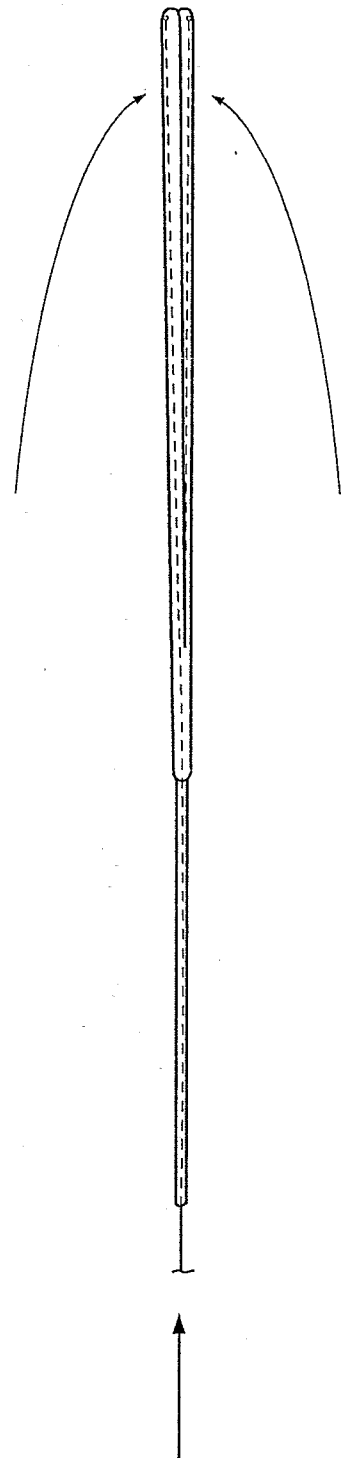


FIG. 9B

11/19

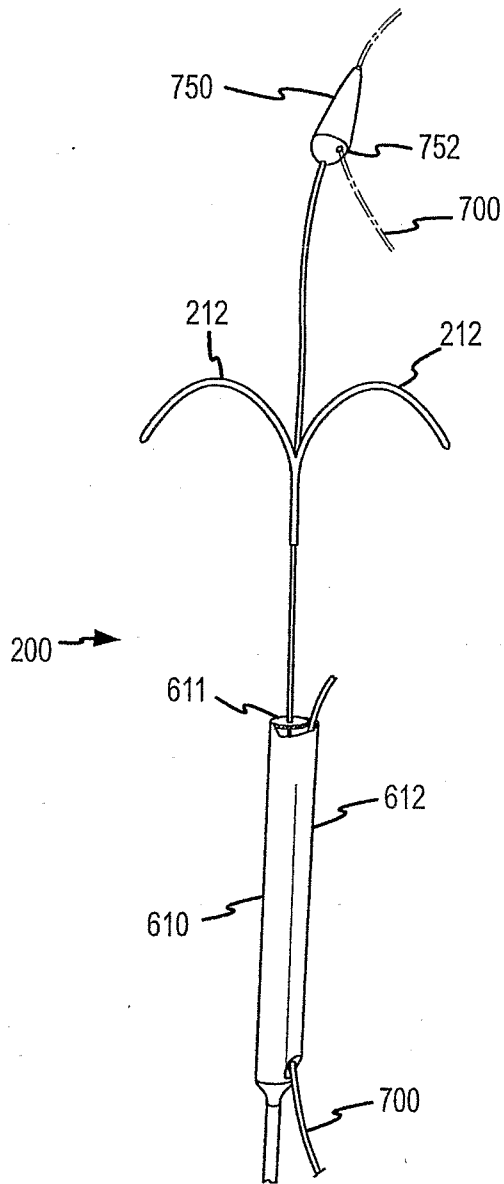


FIG.10A

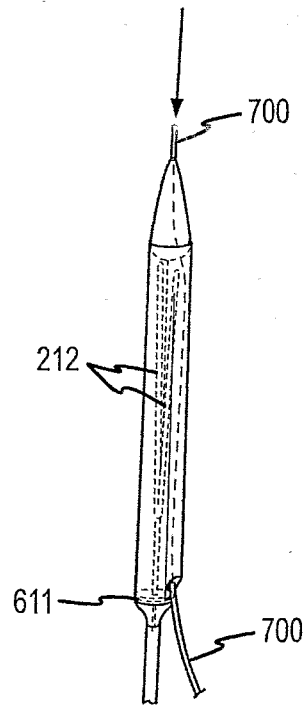


FIG.10B

12/19

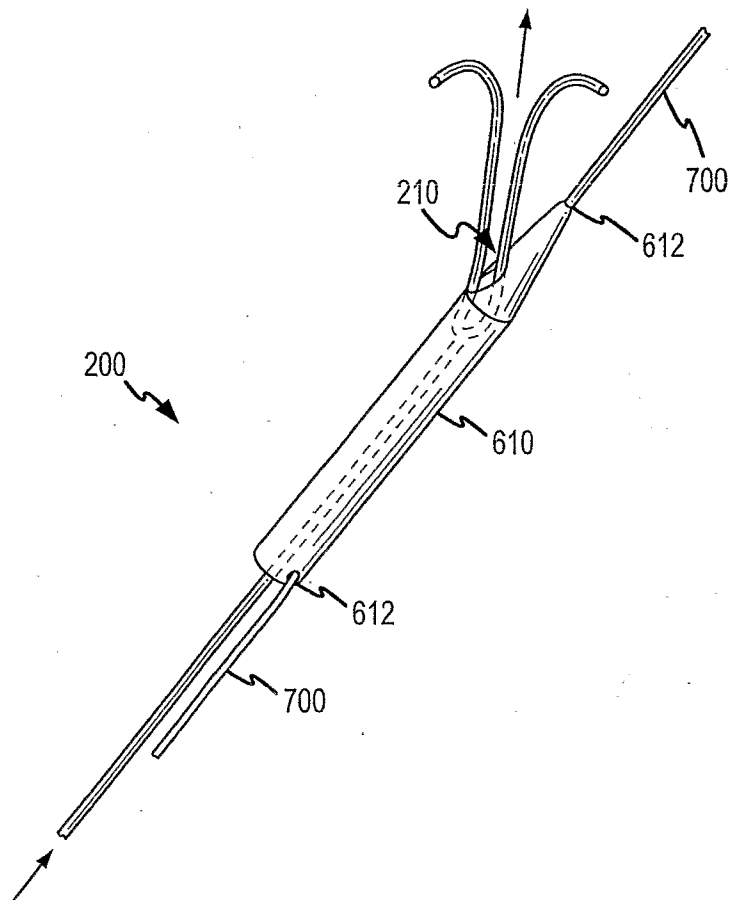


FIG. 11

13/19

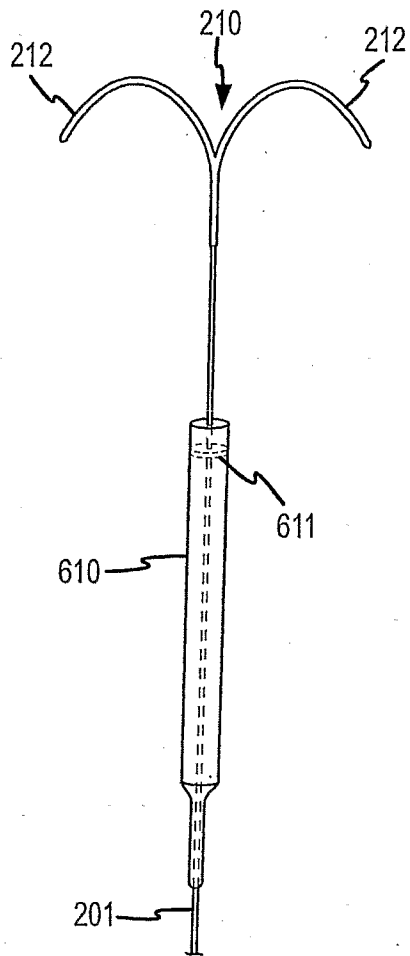


FIG. 12A

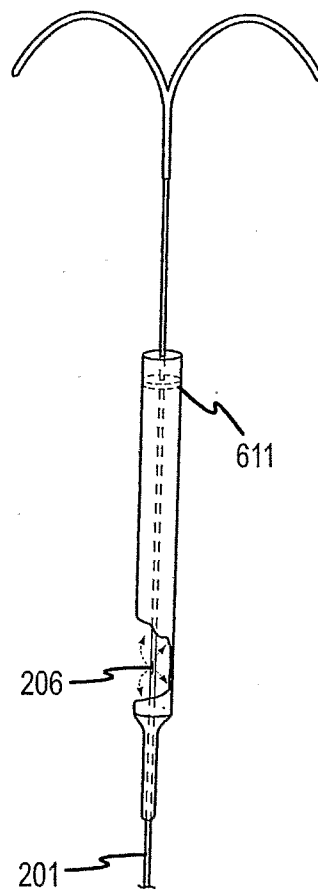


FIG. 12B

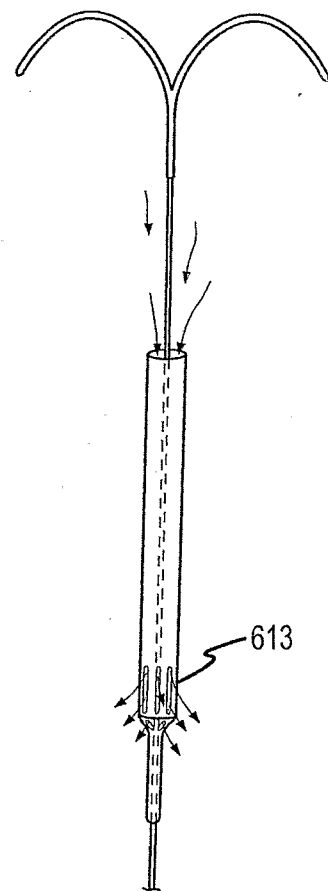


FIG. 12C

14/19

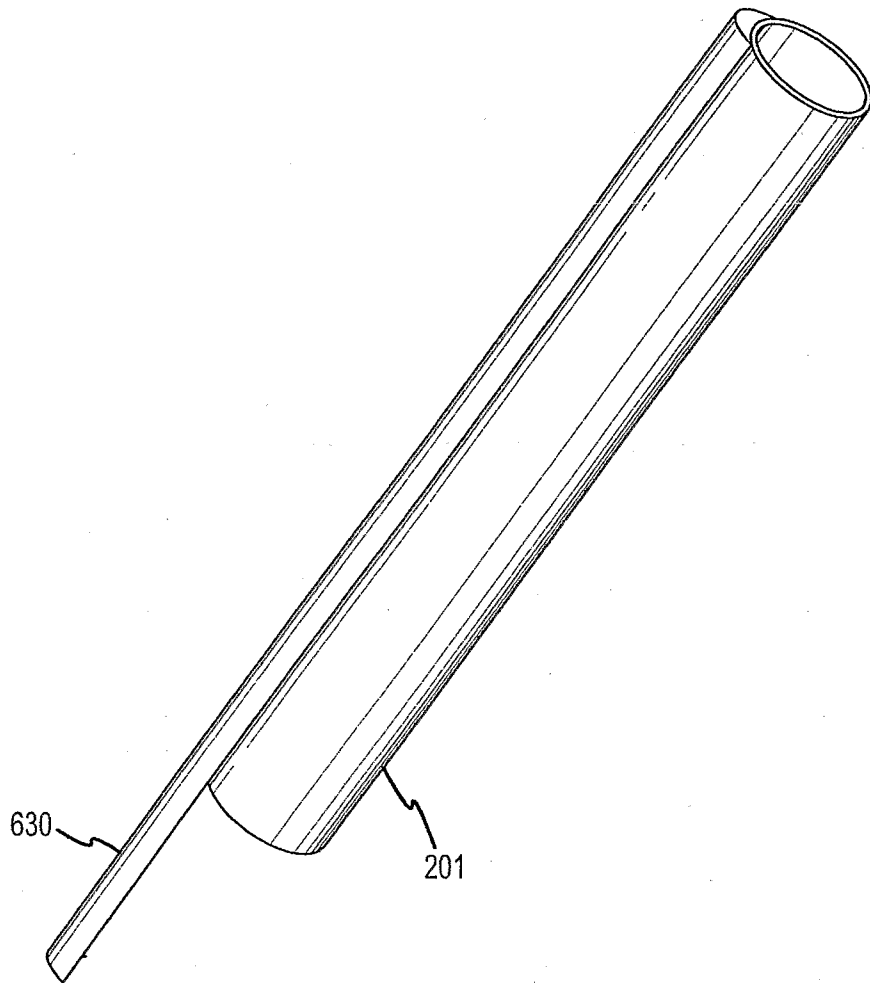


FIG.13

15/19

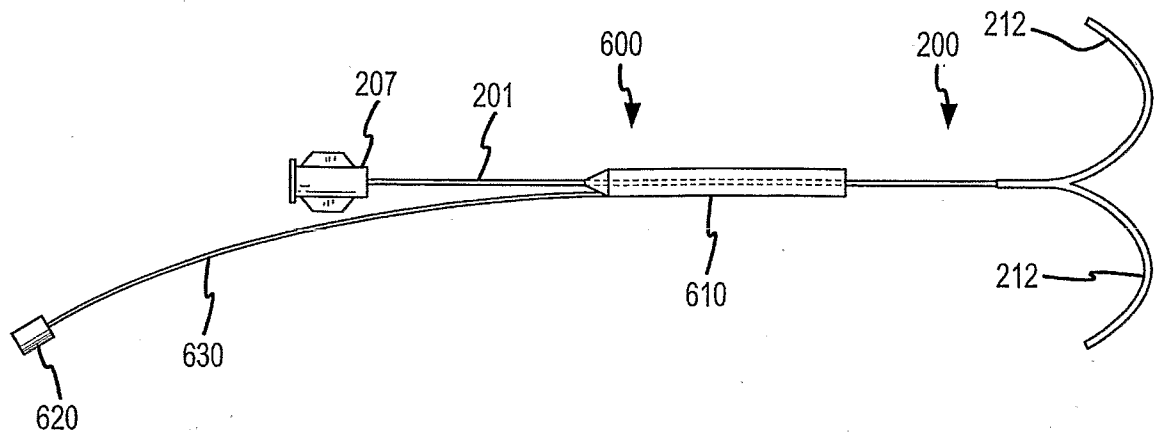


FIG. 14A

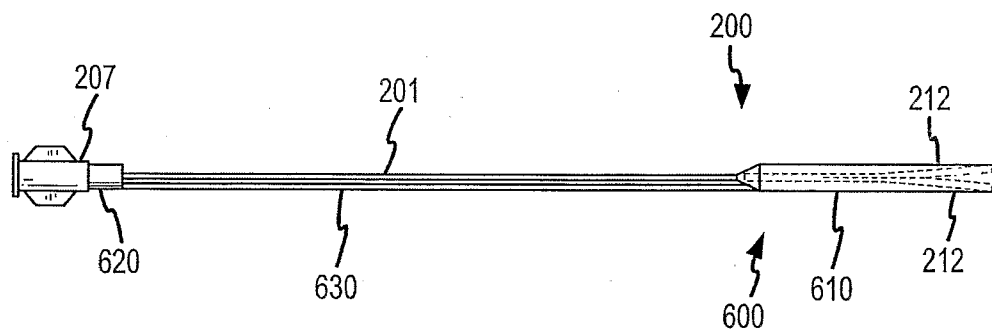


FIG. 14B

16/19

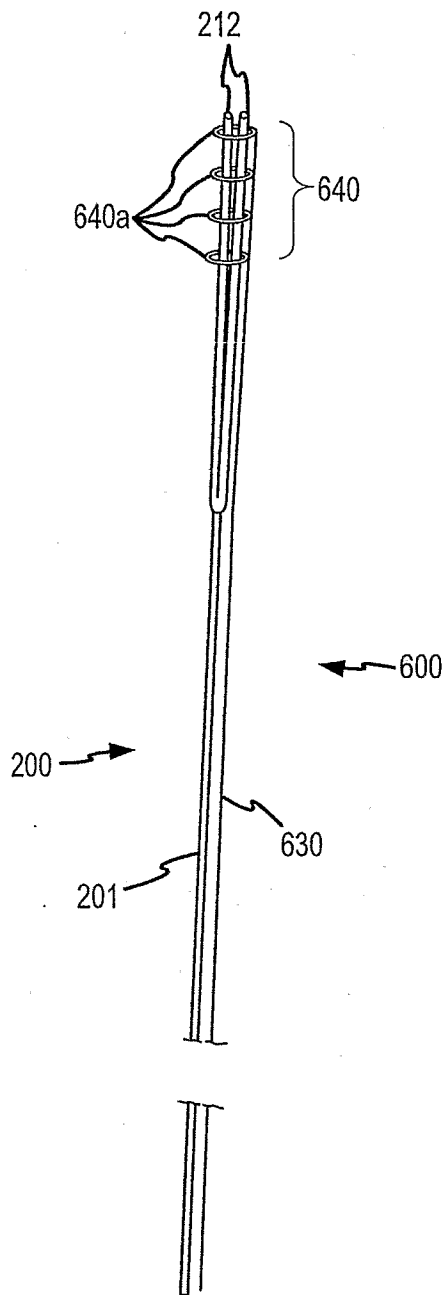


FIG. 15A

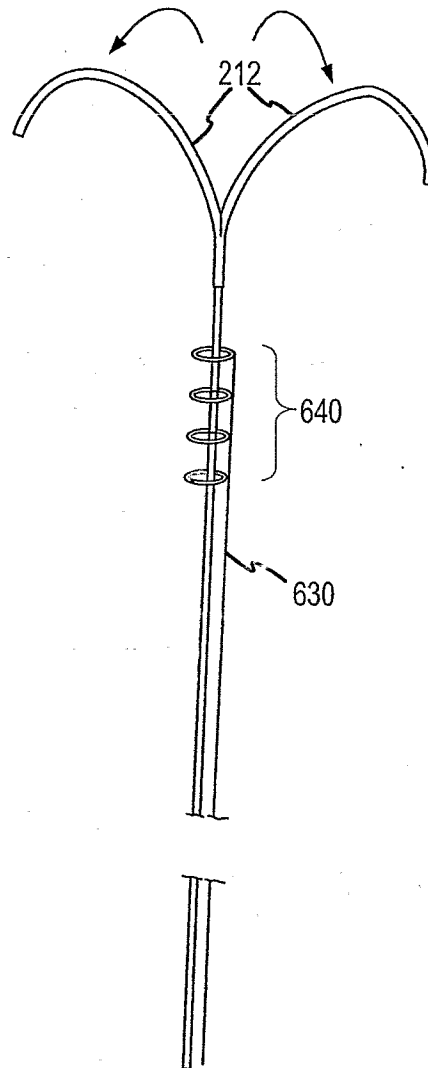


FIG. 15B

17/19

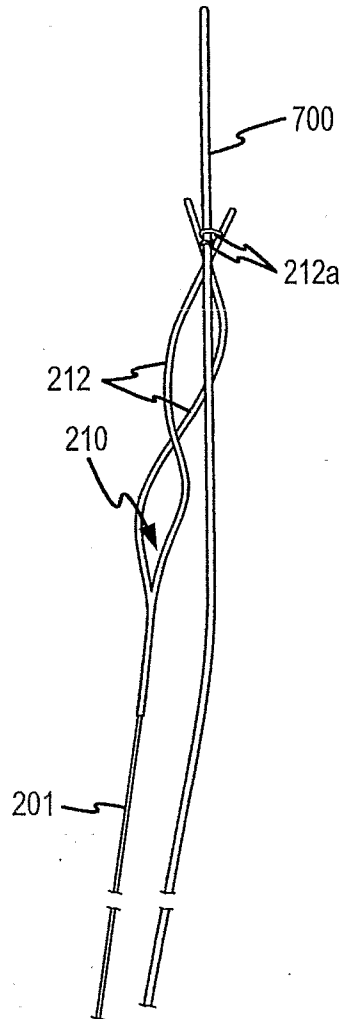


FIG. 16A

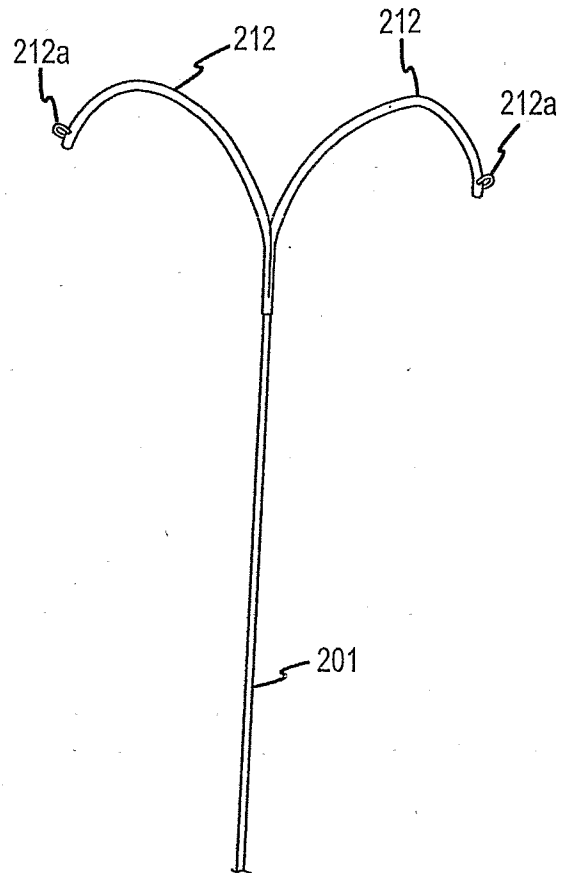


FIG. 16B

18/19

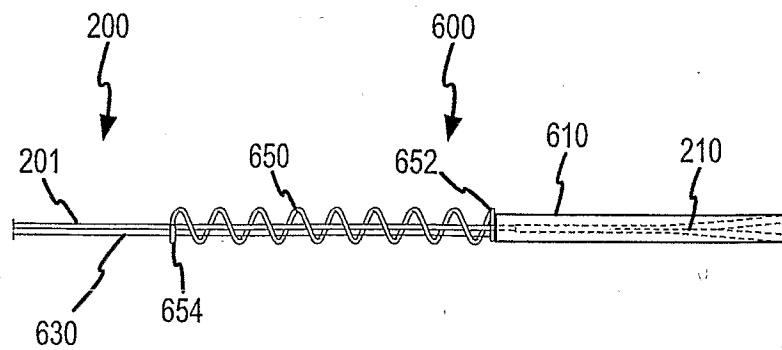


FIG.17A

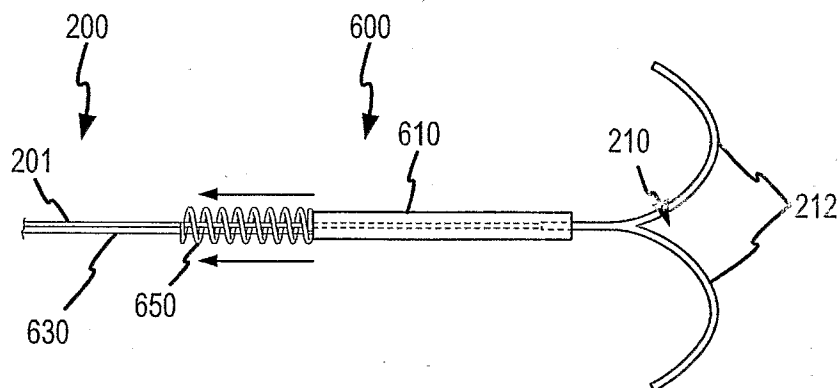


FIG.17B

19/19

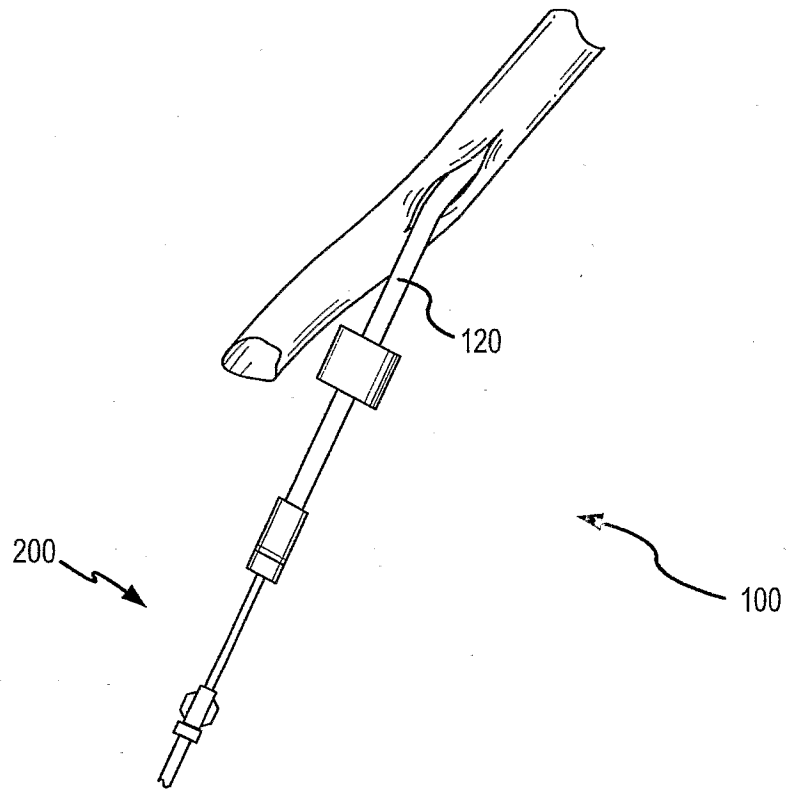


FIG.18