

(12) STANDARD PATENT
(19) AUSTRALIAN PATENT OFFICE

(11) Application No. AU 2009314133 B2

(54) Title
Selective accumulation of energy with or without knowledge of tissue topography

(51) International Patent Classification(s)
A61B 18/14 (2006.01)

(21) Application No: **2009314133** (22) Date of Filing: **2009.11.11**

(87) WIPO No: **WO10/056745**

(30) Priority Data

(31) Number **61/115,344** (32) Date **2008.11.17** (33) Country **US**

(43) Publication Date: **2010.05.20**

(44) Accepted Journal Date: **2015.12.10**

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(56) Related Art
US 4784132
US 2008/0125772

INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) **World Intellectual Property Organization**
International Bureau



(43) **International Publication Date**
20 May 2010 (20.05.2010)

(10) **International Publication Number**
WO 2010/056745 A1

(51) **International Patent Classification:**
A61B 18/14 (2006.01)

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

(21) **International Application Number:**
PCT/US2009/064027

(84) **Designated States** (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, 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, HR, HU, IE, IS, IT, LT, LU, LV, MC, MK, MT, NL, NO, PL, PT, RO, SE, SI, SK, SM, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

(22) **International Filing Date:**
11 November 2009 (11.11.2009)

Published:

(25) **Filing Language:** English

— with international search report (Art. 21(3))

(26) **Publication Language:** English

(30) **Priority Data:**
61/115,344 17 November 2008 (17.11.2008) US

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(54) **Title:** SELECTIVE ACCUMULATION OF ENERGY WITH OR WITHOUT KNOWLEDGE OF TISSUE TOPOGRAPHY

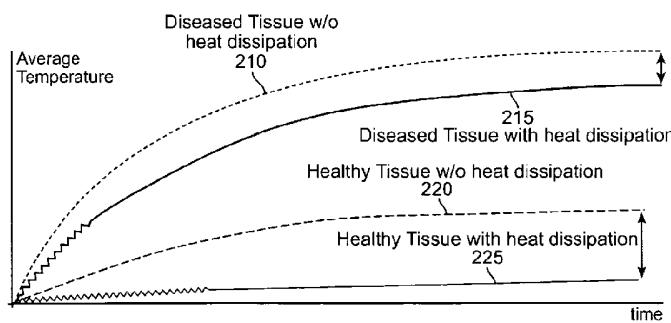


FIG. 5

(57) **Abstract:** Methods and systems for heating a body tissue region adjacent a body lumen with diseased and healthy portions using selective accumulation of energy in the artery tissue with or without knowledge of tissue topography. The method includes positioning an energy delivery portion of a catheter body within the lumen adjacent the body tissue region to be heated, determining a pulse characteristic in response to a thermal property of a first tissue type and applying pulsed energy with the pulse characteristic from the energy delivery portion so as to therapeutically treat the second tissue type within the body tissue region by drawing heat from the first tissue type at a rate that avoids significant thermal damage to the first tissue type while building-up heat in the second tissue type.

WO 2010/056745 A1

SELECTIVE ACCUMULATION OF ENERGY WITH OR WITHOUT KNOWLEDGE OF TISSUE TOPOGRAPHY

CROSS-REFERENCES TO RELATED APPLICATIONS

5 [0001] The present application claims the benefit under 35 USC 119(e) of US Provisional Application No. 61/115,344 filed November 17, 2008; the full disclosure of which is incorporated herein by reference in its entirety.

[0002] This application is related to U.S. Application No. 11/975,474, filed on October 18, 2007, entitled "Inducing Desirable Temperature Effects on Body Tissue"; U.S. Application 10 No. 11/975,383, filed on October 18, 2007, and entitled "System for Inducing Desirable Temperature Effects on Body Tissues", U.S. Patent Application No. 11/122,263, filed on May 3, 2005, entitled "Imaging and Eccentric Atherosclerotic Material Laser remodeling and/or Ablation Catheter", and U.S. Provisional Application No. 61/099,155, filed on September 22, 2008, entitled "Inducing Desirable Temperature Effects On Body Tissue 15 Using Alternate Energy Sources", the full disclosures of which are incorporated herein by reference.

STATEMENT AS TO RIGHTS TO INVENTIONS MADE UNDER FEDERALLY SPONSORED RESEARCH AND DEVELOPMENT

20 [0003] NOT APPLICABLE

BACKGROUND OF THE INVENTION

[0004] The present invention is generally related to medical devices, systems, and methods for heating body tissue. In exemplary embodiments, the invention provides catheter-based 25 treatment for heating body tissue with energy that selectively accumulates in a desired type of tissue, particularly diseased tissue, with or without knowing the tissue topography a-priori.

[0005] Balloon angioplasty and other catheters often are used to open arteries that have been narrowed due to atherosclerotic disease. The trauma associated with balloon dilation can impose significant injury, so that the benefits of balloon dilation may be limited in time. 30 Stents are commonly used to extend the beneficial opening of the blood vessel. Restenosis or

a subsequent narrowing of the body lumen after stenting has occurred in a significant number of cases.

[0006] More recently, drug coated stents (such as Johnson and Johnson's Cypher™ stent, the associated drug comprising Sirolimus™) have demonstrated a markedly reduced 5 restenosis rate, and others are developing and commercializing alternative drug eluting stents. In addition, work has also been initiated with systemic drug delivery (intravenous or oral) which may also improve the procedural angioplasty success rates.

[0007] While drug eluting stents appear to offer significant promise for treatment of atherosclerosis in many patients, there remain many cases where stents either cannot be used 10 or present significant disadvantages. Generally, stenting leaves an implant in the body. Such implants can present risks, including mechanical fatigue, corrosion, and the like, particularly when removal of the implant is difficult and involves invasive surgery. Stenting may have additional disadvantages for treating diffuse artery disease, for treating bifurcations, for treating areas of the body susceptible to crush, and for treating arteries subject to torsion, 15 elongation, and shortening.

[0008] A variety of modified restenosis treatments or restenosis-inhibiting treatment modalities have also been proposed, including intravascular radiation, cryogenic treatments, ultrasound energy, and the like, often in combination with balloon angioplasty and/or stenting. While these and different approaches show varying degrees of promise for 20 decreasing the subsequent degradation in blood flow following angioplasty and stenting, the trauma initially imposed on the tissues by angioplasty remains problematic.

[0009] A number of alternatives to stenting and balloon angioplasty so as to open stenosed arteries have also been proposed. For example, a wide variety of atherectomy devices and techniques have been disclosed and attempted. Despite the disadvantages and limitations of 25 angioplasty and stenting, atherectomy has not gained the widespread use and success rates of dilation-based approaches. More recently, still further disadvantages of dilation have come to light. These include the existence of vulnerable plaque, which can rupture and release materials that may cause myocardial infarction or heart attack.

[0010] In light of the above, it would be advantageous to provide methods and systems for 30 inducing vasodilation in artery tissue and remodeling of the lumens of the body. It would further be desirable to avoid significant cost or complexity while providing structures which

could remodel body lumens without having to resort to the trauma of extreme dilation, and to allow the opening of blood vessels and other body lumens which are not suitable for stenting.

BRIEF SUMMARY

[0010a] It is an object of the present disclosure to substantially overcome, or at least ameliorate, at least one disadvantage of present arrangements.

[0011] In a first aspect, the present disclosure provides a method for heating a body tissue region adjacent a body lumen, the region having both a first tissue type and a second tissue type. The method includes positioning an energy delivery portion of a catheter body within the lumen adjacent the body tissue region to be heated, determining a pulse characteristic in response to a thermal property of the first tissue type and applying pulsed energy with the pulse characteristic from the energy delivery portion so as to therapeutically treat the second tissue type within the body tissue region by drawing heat from the first tissue type at a rate that avoids significant thermal damage to the first tissue type while building-up heat in the second tissue type.

[0012] In another aspect, the present disclosure provides a catheter system for heating a body tissue region adjacent a body lumen, the region having both a first tissue type and a second tissue type. The system includes an elongate flexible catheter body having a proximal end and a distal end with an axis therebetween, an energy delivery portion proximate the distal end, an energy source coupled to the energy delivery portion and a processor coupled to the energy source, the processor configured to control a pulse characteristic of pulsed energy transmitted from the energy source to the energy delivery portion so as to therapeutically treat the second tissue type within the body tissue region by drawing heat from the first tissue type at a rate that avoids significant thermal damage to the first tissue type while building-up heat in the second tissue type.

[0013] In some aspects, the pulsed energy preferentially heats the second tissue type more than the first tissue type, the preferential heating induced at least in part by one or more of the following group of characteristics of the second tissue type: lower thermal conduction than a thermal conduction of the first tissue type, lower specific heat capacity than a specific heat capacity of the first tissue type, less innate blood perfusion than an innate blood perfusion of the first tissue type, and/or larger distance away from well-perfused areas than a distance between well-perfused areas and the first tissue type.

[0013a] In a further aspect of the present disclosure there is provided a method for heating a body tissue region adjacent a body lumen, the region having both a first tissue type and a second tissue type, the method comprising: positioning an energy delivery portion of a catheter within

the lumen adjacent the body tissue region to be heated, wherein the energy deliver portion comprises a radially expandable structure having a plurality of electrodes thereon; expanding the radially expandable structure with the body lumen so as to radially engage the body lumen with the plurality of electrodes, wherein the plurality of electrodes engaged against the body lumen complete a plurality of circuits, wherein at least some of the circuits include a first tissue type and a second tissue type while other circuits include the first tissue type without the second tissue type; determining a pulse characteristic in response to a thermal property of the first tissue type; applying pulsed electrical energy with the pulse characteristic from the energy delivery portion through the plurality of circuits without identifying which of the plurality of circuits include the second tissue type so that both the first tissue type and the second tissue type are heated by thermal conduction, and while the pulsed electrical energy is applied to the other circuits at a pulsed rate that avoids thermal damage to the first tissue type, applying the same pulsed energy to the at least some circuits at the pulsed rate so as to therapeutically treat the second tissue type with heat; and controlling the pulse characteristic in response to differing values of the thermal property of the first tissue type during application of pulsed electrical energy.

[0013b] In a still further aspect, the present invention provides a catheter system for heating a body tissue region adjacent a body lumen, the region having both a first tissue and a second tissue type, the system comprising: an elongate flexible catheter body having a proximal end and a distal end with an axis therebetween; the catheter body having a radially expandable structure near the distal end; an energy delivery portion proximate the distal end; wherein the energy deliver portion comprises a plurality of electrodes positioned about the radially expandable structure such that when expanded within the body lumen the plurality of electrodes radially engage the body lumen, wherein the plurality of electrodes engaged against the body lumen complete a plurality of circuits, wherein at least some of the circuits include a first tissue type and a second tissue type while other circuits include the first tissue type without the second tissue type; an energy source coupled to the energy delivery portion; and a processor coupled to the energy source, the processor configured to control a pulse characteristic of pulsed energy transmitted from the energy source to each of the plurality of circuits without identifying which of the plurality of circuits include the second tissue type so that both the first tissue type and the second tissue type are heated by the thermal conduction, wherein the energy delivery portion is configured to apply the pulsed electrical energy to the other circuits at a pulsed rate that avoids thermal damage to the first tissue type, and to apply the same pulsed energy to the at least some circuits at the pulsed rate so as to therapeutically treat the second tissue type with heat.

[0014] In some aspects, most group of characteristics cause heat to be drawn from the first tissue type at a rate that avoids significant thermal damage to the first tissue type, while allowing heat to build up in the second tissue type.

[0015] In some aspects, the pulse energy is delivered at an average rate of 0.25 to 5 watts to the body tissue region.

[0016] In some aspects, each pulse of the pulsed energy provides between 4 to 45 Joules to the body tissue region.

[0017] In some aspects, wherein an average rate of pulsed energy delivered to the body tissue region is between about 0.1 and 10.0 times the rate of energy dissipation by the first tissue type.

[0018] In some aspects, a time period between pulses allows the first tissue type to dissipate its heat adequately to avoid thermal damage to the first tissue type by the pulsed energy.

[0019] In some aspects, the time period between pulses of energy is between 0.1 to 180 seconds.

[0020] In some aspects, a thermal time constant for the first tissue type is approximately how long it will take a discrete volume of that given tissue to lose 63% of its heat while undergoing an exponential decay.

[0021] In some aspects, the pulsed energy is delivered to the body tissue region over a duration of at least several first tissue type time constants, wherein the thermal time constant of the first tissue is proportional to a thermal conductivity of the first tissue type.

[0022] In some aspects, the thermal conductivity of the second tissue type is twice that of the first tissue type.

[0023] In some aspects, the energy source comprises a radio frequency (RF) energy source.

[0024] In some aspects, the energy delivery portion comprises a radially expandable structure engaging a plurality of electrode surfaces against the body lumen, wherein the plurality of electrode surfaces against the body lumen complete a plurality of circuits comprising the first tissue type and the second tissue type and pulsed energy is delivered to the plurality of circuits.

- [0025] In some aspects, the energy source comprises a laser energy source.
- [0026] In some aspects, the energy delivery portion comprises at least one radially oriented window coupled to at least one optical conduit extending between the proximal end of the catheter body and at least one window for transmission of pulsed laser energy to the body tissue region.
- [0027] In some aspects, the energy source comprises an ultrasound transmitter configured to deliver pulsed ultrasound energy to the body tissue region.
- [0028] In some aspects, the energy source comprises a microwave energy source including at least one microwave antenna configured to deliver pulsed microwave energy to the body tissue region.

BRIEF DESCRIPTION OF THE DRAWINGS

- [0029] FIG. 1 schematically illustrates one embodiment of a catheter system for heating artery tissue.
- [0030] FIG. 2 illustrates one embodiment of a balloon for use in the catheter system of FIG. 1.
- [0031] FIG. 3 shows what a temperature vs. time profile for heating both healthy tissue and diseased tissue at a constant pulsed rate of heat or energy.
- [0032] FIG. 4 shows another temperature vs. time profile for tissue having both healthy tissue and diseased tissue in which the heat or energy is applied at different rates.
- [0033] FIG. 5 shows heat dissipation's effect on temperature.
- [0034] FIG. 6 schematically illustrates one embodiment of balloon catheter system for use for bipolar treatment of tissue in a leg.
- [0035] FIG. 7 schematically illustrates one embodiment of balloon catheter system for use for monopolar treatment of tissue in a leg.
- [0036] FIGS. 8A-8C illustrate a method of using a balloon catheter system treating artery tissue.

DETAILED DESCRIPTION OF THE INVENTION

[0037] Many therapies have been developed to replace or improve upon traditional balloon angioplasty and stents. Many of the devices described in the BACKGROUND OF THE INVENTION either cut, ablate, or vaporize diseased tissue in an artery. For example, laser 5 devices vaporize plaque and flush it downstream. Atherectomy devices excise plaque and suck it out of the body. Cutting balloons incise the artery wall, damaging the tissue.

[0038] It would be advantageous to provide systems and devices that do not cut, ablate, or vaporize. Three modalities of treatment avoid these drawbacks, they include: cooling the tissue; non-ablative forms of direct molecular denaturing; and non-ablative heating. Cooling 10 has been implemented using devices such as Boston Scientific's Cryo-cath. Direct molecular denaturing can be achieved with radiation -- gamma rays, for instance. The present invention is directed to the remaining modality, non-ablative heating.

[0039] Some embodiments of the present invention generally provide devices, systems, and methods for heating artery tissue with diseased portions using selective accumulation of 15 energy in the artery tissue with or without knowledge of tissue topography, discussed in more detail below. The invention will be particularly useful for remodeling materials along a partially occluded artery in order to open the artery lumen and increase blood flow. The devices, systems, and methods disclosed herein may be used in any artery, for example, the femoral, popliteal, coronary and/or carotid arteries. Devices for heating artery tissue have 20 been disclosed in co-pending U.S Patent Application Nos. 11/975,474, 11/975,383, 11/122,263 and U.S. Provisional Application No. 61/099,155, the full disclosures of which are incorporated herein by reference.

[0040] While the disclosure focuses on the use of the technology in the vasculature, the 25 technology would also be useful for any luminal obstruction. Other anatomical structures in which the present invention may be used are the esophagus, the oral cavity, the nasopharyngeal cavity, the auditory tube and tympanic cavity, the sinus of the brain, the arterial system, the venous system, the heart, the larynx, the trachea, the bronchus, the stomach, the duodenum, the ileum, the colon, the rectum, the bladder, the ureter, the ejaculatory duct, the vas deferens, the urethra, the uterine cavity, the vaginal canal, and the 30 cervical canal.

[0041] Some embodiments of the system will be able to treat tissue by gentle heating in combination with dilation of the artery. The heating of the vessel wall may be done before,

during, and/or after dilation with the balloon, with dilation pressures which are at or significantly lower than standard, unheated angioplasty dilation pressures. For example, where balloon inflation pressures of 10-16 atmospheres may be appropriate for standard angioplasty dilation of a particular lesion, modified dilation treatments combined with gentle

5 heating described herein may employ from 10-16 atmospheres or may be effected with pressures of 6 atmospheres or less, and possibly as low as 1 to 2 atmospheres. In case of calcification in the artery plaque, it may be more difficult to remodel and open the diseased artery, so the catheter may use a standard angioplasty balloon in combination with ultrasonic energy to break down the calcium and remodel and open the lumen.

10 [0042] The gentle heating energy added before, during, and/or after dilation of a blood vessel may increase dilation effectiveness while lowering complications. In some embodiments, such controlled heating with balloon dilatation may exhibit a reduction in recoil, providing at least some of the benefits of a stent-like expansion without the disadvantages of an implant. Benefits of gentle heating may be enhanced (and/or complications inhibited) by limiting heating of the adventitial layer below a deleterious response threshold. In many cases, such heating of the intima and/or media may be provided using heating times of less than about 10 seconds, often being less than 3 (or even 2) seconds. In other cases, very low power may be used for longer durations.

15 [0043] Heating of the body tissue involves the application of energy, typically in the form of RF, microwave and/or ultrasound energy, and the like. This energy will be controlled so as to limit a temperature of the body tissue. In some embodiments, the body tissue temperature range is from about 50°C to about 90°C, depending on the energy applied and tissue type.

20 [0044] While the methods and devices described herein are not selective in tissue treatment of the blood vessel, the devices can be used for treatment of both concentric and eccentric atherosclerosis. This non selective treatment is a particular advantage because atherosclerosis may be eccentric relative to an axis of the blood vessel over 50% of the time, possibly in as much as (or even more than) 75% of cases.

25 [0045] While the present invention may be used in combination with stenting, the present invention is particularly well suited for increasing the open diameter of blood vessels in which stenting is not a viable option. Potential applications include treatment of diffuse disease, in which atherosclerosis is spread along a significant length of an artery rather than

being localized in one area. The present invention may also find advantageous use for treatment of tortuous, sharply-curved vessels, as no stent need be advanced into or expanded within the sharp bends of many blood vessel. Still further advantageous applications include treatment along bifurcations (where side branch blockage may be an issue) and in the 5 peripheral extremities such as the legs, feet, and arms (where crushing and/or stent fracture failure may be problematic).

[0046] The disclosed system includes at least an energy source and a catheter with an energy delivery portion configured for heating a body tissue region adjacent a body lumen, the region having both a first tissue type, such as healthy tissue, and a second tissue type, 10 such as diseased tissue. The catheter may be similar to a balloon catheter commonly used to treat artery disease today, except for the addition of electrodes used for heating. Other embodiments may use transducers disposed on the balloon to apply ultrasound heating to the vessel wall or microwave antennas disposed on the balloon to apply microwave heating to the vessel wall. The energy source may be coupled to a processor to control a pulse 15 characteristic of pulsed energy transmitted from the energy source to the energy delivery portion so as to therapeutically treat the tissue within the body treatment region without causing sufficient thermal damage to the body treatment region so as to induce a long-term occlusive response.

[0047] In use, the energy delivery portion of a catheter body is positioned within the lumen 20 adjacent the body tissue region to be heated. A pulse characteristic is determined in response to a thermal property of the first tissue type, and the pulsed energy is applied with the pulse characteristic from the energy delivery portion so as to therapeutically treat the body tissue region.

[0048] Fig. 1 shows one embodiment of a catheter system 10 for heating artery tissue. The 25 catheter system 10 includes a balloon catheter 12 having a catheter body 14 with a proximal end 16 and a distal end 18. Catheter body 14 is flexible and defines a catheter axis 15, and may include one or more lumens, such as a guidewire lumen 22 and an inflation lumen 24 (see Fig. 2). Still further lumens may be provided if desired for other treatments or 30 applications, such as perfusion, fluid delivery, imaging, or the like. Catheter 12 includes an inflatable balloon 20 adjacent distal end 18 and a housing 29 adjacent proximal end 16. Housing 29 includes a first connector 26 in communication with guidewire lumen 22 and a second connector 28 in fluid communication with inflation lumen 24. Inflation lumen 24

extends between balloon 20 and second connector 28. Both first and second connectors 26, 28 may optionally comprise a standard connector, such as a Luer-Loc™ connector. A distal tip may include an integral tip valve to allow passage of guidewires, and the like.

[0049] Housing 29 also accommodates an electrical connector 38. Connector 38 includes a 5 plurality of electrical connections, each electrically coupled to electrodes 34 via conductors 36. This allows the electrodes 34 to be easily energized, the electrodes often being energized by a controller 40 and energy source 42, such as bipolar or monopolar RF energy, microwave energy, ultrasound energy, or other suitable energy sources. In one embodiment, electrical connector 38 is coupled to an RF generator via a controller 40, with controller 40 allowing 10 energy to be selectively directed to electrodes 38. When monopolar RF energy is employed, patient ground may (for example) be provided by an external electrode or an electrode on catheter body 14. .

[0050] In some embodiments, controller 40 may include a processor or be coupled to a processor to control or record treatment. The processor will typically comprise computer 15 hardware and/or software, often including one or more programmable processor unit running machine readable program instructions or code for implementing some or all of one or more of the methods described herein. The code will often be embodied in a tangible media such as a memory (optionally a read only memory, a random access memory, a non-volatile memory, or the like) and/or a recording media (such as a floppy disk, a hard drive, a CD, a 20 DVD, a non-volatile solid-state memory card, or the like). The code and/or associated data and signals may also be transmitted to or from the processor via a network connection (such as a wireless network, an Ethernet, an internet, an intranet, or the like), and some or all of the code may also be transmitted between components of catheter system 10 and within 25 processor via one or more bus, and appropriate standard or proprietary communications cards, connectors, cables, and the like will often be included in the processor. Processor will often be configured to perform the calculations and signal transmission steps described herein at least in part by programming the processor with the software code, which may be written as a single program, a series of separate subroutines or related programs, or the like. The processor may comprise standard or proprietary digital and/or analog signal processing 30 hardware, software, and/or firmware, and will typically have sufficient processing power to perform the calculations described herein during treatment of the patient, the processor optionally comprising a personal computer, a notebook computer, a tablet computer, a proprietary processing unit, or a combination thereof. Standard or proprietary input devices

(such as a mouse, keyboard, touchscreen, joystick, etc.) and output devices (such as a printer, speakers, display, etc.) associated with modern computer systems may also be included, and processors having a plurality of processing units (or even separate computers) may be employed in a wide range of centralized or distributed data processing architectures.

5 [0051] Balloon 20 is illustrated in more detail in Fig. 2. Balloon 20 generally includes a proximal portion 30 coupled to inflation lumen 24 and a distal portion 32 coupled to guidewire lumen 22. Balloon 20 expands radially when inflated with a fluid or a gas. In some embodiments, the fluid or gas may be non-conductive and/ cooled. In some embodiments, balloon 20 may be a low pressure balloon pressurized to contact the artery 10 tissue. In other embodiments, balloon 20 is an angioplasty balloon capable of higher pressure to both heat the artery tissue and expand the artery lumen. Balloon 20 may comprise a compliant or non-compliant balloon having helical folds to facilitate reconfiguring the balloon from a radially expanded, inflated configuration to a low profile configuration, particularly for removal after use.

15 [0052] Electrodes 34 are mounted on a surface of balloon 20, with associated conductors 36 extending proximally from the electrodes. Electrodes 34 may be arranged in many different patterns or arrays on balloon 20. The system may be used for monopolar or bipolar application of energy. For delivery of monopolar energy, a ground electrode is used, either on the catheter shaft, or on the patients skin, such as a ground electrode pad. For delivery of 20 bipolar energy, adjacent electrodes are axially offset to allow bipolar energy to be directed between adjacent circumferential (axially offset) electrodes. In other embodiments, electrodes may be arranged in bands around the balloon to allow bipolar energy to be directed between adjacent distal and proximal electrodes. In some embodiments, the electrodes 34 may be positioned internal of balloon 20. Electrodes 34 may be arranged in many different 25 patterns or arrays on balloon 20. As mentioned previously, other embodiments may include ultrasound transducers or microwave antennas mounted on a surface of balloon 20.

[0053] The embodiments disclosed herein revolve around the concept of selective accumulation of energy in artery tissue with or without knowledge of tissue topography. This is accomplished by taking advantage of the differences in tissue properties between healthy 30 tissue and diseased tissue. The preferential heating of the tissues with different properties can be accomplished without knowing the location or type the different tissues. If a first tissue type has a better thermal conductivity (k) than a second tissue type, it will conduct heat away

more rapidly. If the second tissue type has a lower specific heat capacity (c_p) than the first tissue type, its temperature will increase more given the same amount of energy applied to the same mass (and volume, assuming relatively similar tissue density). If the first tissue type has denser vasculature, or is reliably in closer proximity to well-perfused areas, it will 5 conduct heat away more rapidly than the second tissue type.

[0054] This present invention allows preferentially heating of a type of tissue that has one or more of the following characteristics:

- Relatively poor (lower) thermal conduction,
- Lower specific heat capacity,
- 10 • Less innate blood perfusion, and/or
- Relatively larger distance away from well-perfused areas.

[0055] In the case of diseased tissue, all of the above characteristics apply. The disease is generally comprised of lipidic fat-like diseased tissue and/or fibrous collagen-like tissue. Both of these tissues have a lower specific heat capacity and lower thermal conductivity than 15 healthy vascular tissue, particularly media. Healthy vascular tissue also has more microvasculature, and is in closer proximity to well-perfused tissue, therefore healthy tissue can sink heat away more effectively, without the heat flux “backing up” like a traffic jam.

[0056] The key to taking advantage of this discrepancy in tissue properties is the following: Heat is applied or generated within the tissue at a rate commensurate with the thermal time 20 constants of the tissues involved, possibly using a pulse width modulation (PWM) approach. During each “pulse”, the same quantity of energy is delivered or generated regardless of the tissue type. Tissue topography may be unknown, therefore the same “treatment” is applied everywhere. However, in the diseased tissue, the temperature profile will be higher 25 everywhere because it has a lower heat capacity. The diseased tissue will also retain its heat longer because it has lower thermal conductivity and therefore a longer thermal time constant. Thus, the healthy tissue will heat up slower and cool off faster, which is exactly what is desired for this particular application. The goal is to heat diseased tissue while not heating healthy tissue. The amount of time between pulses can then be tailored to allow the 30 healthy tissue to cool and dissipate its heat adequately so as not to exceed a particular amount of thermal damage the healthy tissue.

[0057] FIG. 3 shows what a temperature vs. time profile for heating both healthy tissue 110 and diseased tissue 120 at a constant pulsed rate of heat or energy. The diseased tissue

temperature will rise faster and cool slower than the healthy tissue, therefore accumulating heat and receiving treatment significantly more than healthy tissue.

[0058] FIG. 4 shows another a temperature vs. time profile for tissue having both healthy tissue and diseased tissue in which the heat or energy is applied at different rates. The figure 5 shows it is possible to initially apply or generate heat faster (at the beginning of the treatment) in order to ramp up the tissue temperature more quickly. This could be done with power modulation or PWM with a variable pulse width. The purpose of this would be to heat the tissue quickly and then allow the healthy tissue to dissipate heat between pulses, which would decrease the total treatment time required to achieve (and possibly soak at) a certain 10 temperature.

[0059] TIME CONSTANT:

$$t = RC = \left(\frac{l}{n} k \right) (mc_p), \text{ where healthy } t \approx 7\text{s and diseased } t \approx 14\text{s.}$$

[0060] HEAT CAPACITY:

Media $c_p \approx 3.9$

15 Adventitia $c_p \approx 3.1$

Fat $c_p \approx 2.4$

[0061] THERMAL CONDUCTIVITY:

Media $k \approx .59$

Adventitia $k \approx .49$

20 Fat $k \approx .20$

[0062] The “TIME CONSTANT” above is the thermal time constant. This is approximately how long it will take a discrete volume of that given tissue to lose 63% of its heat (and therefore temperature) while undergoing an exponential decay. The thermal time constant is directly proportional to the thermal conductivity. Therefore, because there is 25 approximately a 2:1 ratio between the thermal conductivity of fatty diseased tissue and healthy media tissue, there is also a 2:1 ratio between their time constants.

[0063] The thermal time constant for healthy media tissue was estimated to be 7 seconds based upon a treatment volume of 32 mm³, a thermal conductivity of 0.5 W/m/K, a thermal

front cross-sectional area of 32 mm², and a specific heat capacity of 3.6 J/g/K. In this case, one would want to deliver an appropriate therapeutic dose of energy over a duration of at least several time constants, e.g. 35 seconds, 70 seconds, etc.

[0064] FIG. 5 shows heat dissipation's effect on temperature versus time. Because thermal 5 dissipation will happen according to an exponential decay, the rate of cooling will vary nonlinearly with time. So, the period has to be selected appropriately in conjunction with the power and duty cycle. The power and the heat capacity, neglecting heat dissipation, will determine the rate of temperature increase during heating. The combination of power, duty cycle, heat capacity, and the rate of continuous heat dissipation, will determine the average 10 rate of temperature change. Diseased tissue 210 and healthy tissue 220 both have higher average temperatures without heat dissipation than diseased tissue 215 and healthy tissue 225 with heat dissipation.

[0065] The important thing to note about FIG. 5 is that the average power (average energy 15 per time) has to be low enough that the rate of heat dissipation is significant in comparison. In other words, if we were to assume that healthy vessel tissue in a certain geometry can effectively dissipate energy at a rate of 2 Joules per second (2 Watts), then the average rate of energy application needs to be on this order of magnitude as well, in order for the effect to be significant.

[0066] This is the main crux of the invention – applying energy at an average rate 20 commensurate with the thermal time constants involved such that significant differences in the tissues' thermal properties can be leveraged to our advantage.

[0067] Fig. 6 schematically illustrates bipolar treatment of tissue with the system 10. Balloon 20 having electrode pairs 34A and 34B is positioned within an artery lumen having fatty disease/necrotic core 48, fibrous disease/fibrous cap 44, healthy tissue 45. Treatment is 25 done to healthy tissue 45 and the fatty disease/necrotic core 48, fibrous disease/fibrous cap 44 by pulsed energy between electrode pairs 34A and 34B. The electrode pairs may be any electrode pairs on the balloon, for example, in some embodiments, the electrode pairs may be 34A and 34C, or 34A and 34D, or any combination of 34A-34D. This arrangement creates an energy path 50 through the tissue that delivers energy or heat ("tissue remodeling energy") 30 to the artery tissue between the electrode pairs ("remodeling zones"). Using different combinations of electrode pairs may reduce or eliminate gaps between the remodeling zones by using overlapping pairs. Using electrode pairs with bipolar energy may avoid some

potential issues of the monopolar approach. Diseased artery tissue 48 has a higher electrical resistivity than healthy artery tissue 45. By using pairs of electrodes 34A, 34B in a bipolar system, the energy path 50 will go through the healthy tissue, diseased tissue, or a combination of both healthy and diseased tissues between the electrode pairs. Any number of 5 electrode pairs may be used in different patterns or arrays.

[0068] Fig. 7 schematically illustrates bipolar treatment of tissue with the system 10. Balloon 20 having electrode pairs 34A and 34B is positioned within an artery lumen having fatty disease/necrotic core 48, fibrous disease/fibrous cap 44, healthy tissue 45 and one or more electrical grounds (not shown) are used, such as positioned on the patients skin. When 10 power is applied to the multiple monopolar electrodes 34 arranged around the circumference of the artery lumen, energy 54 is directed radially outward through the artery wall and treats both diseased and healthy artery tissue.

[0069] The use of catheter system 10 for treating tissue by pulsed energy can be understood with reference to Figs. 8A-8C. As seen in Fig. 8A, accessing of a treatment site 15 will often involve advancing a guidewire 56 within a blood vessel 58 at a target region of diseased tissue 48. Location of balloon 20 may be facilitated by radiopaque markers or by radiopaque structure (or corresponding radiopaque markers placed on or near) balloon 20, and/or by the use of radiopaque electrodes 34. Guidewire 56 may be positioned under fluoroscopic (or other) imaging.

20 [0070] Catheter 12 is advanced distally over guidewire 56 and positioned adjacent to atherosclerotic material 48. Balloon 20 expands radially within the lumen of the blood vessel so that electrodes 34, or electrodes 34A and 34B, radially engage artery tissue. As diseased tissue 48 may be distributed eccentrically about catheter 12, electrodes 34 may engage diseased tissue 48, healthy tissue 60, or a combination of both tissues.

25 [0071] As discussed above, electrodes 34 are positioned circumferentially around the balloon 20. Pulsed energy is directed to electrodes 34, or adjacent pairs of electrodes 34A and 34B, treating both diseased tissue 48 and the healthy tissue 60. The controller 40 may provide pulsed energy to energize the electrodes with about 0.25 to 5 Watts average power for 0.1 to 180 seconds, or with about 4 to 45 Joules. The power and duration are calibrated to 30 be less than enough to cause severe damage, and particularly less than enough to ablate tissue within a blood vessel.

[0072] Referring now to Fig. 8C, as described above, balloon 20 may be an angioplasty balloon that combines heating with opening the artery lumen, such that the heat is preferentially delivered to the diseased tissue 48, for example to mildly heat a cap structure (to induce thickening of the cap and make the plaque less vulnerable to rupture) and/or heat a 5 lipid-rich pool of the vulnerable plaque (so as to remodel, denature, melt, shrink, and/or redistribute the lipid-rich pool), while the healthy tissue 60 dissipates the heat without damage.

[0073] In some embodiments, balloon 20 may be repeatedly contracted, axial movement of the catheter 12 employed to reposition balloon 20, with subsequent expansion of balloon 20 10 at each of a plurality of treatment locations along diseased tissue.

[0074] While generally described herein with reference to the vasculature, embodiments of the catheter devices, systems, and methods described herein may also find applications in the lumens of other vessels of the human anatomy having tissue types with different tissue properties, as discussed above. The anatomical structure into which the catheter is placed 15 may be for example, the esophagus, the oral cavity, the nasopharyngeal cavity, the auditory tube and tympanic cavity, the sinus of the brain, the larynx, the trachea, the bronchus, the stomach, the duodenum, the ileum, the colon, the rectum, the bladder, the ureter, the ejaculatory duct, the vas deferens, the urethra, the uterine cavity, the vaginal canal, and the cervical canal, as well as the arterial system, the venous system, and/or the heart.

20 [0075] While the exemplary embodiments have been described in some detail, by way of example and for clarity of understanding, those of skill in the art will recognize that a variety of modifications, adaptations, and changes may be employed. Hence, the scope of the present invention should be limited solely by the claims.

CLAIMS:

1. A method for heating a body tissue region adjacent a body lumen, the region having both a first tissue type and a second tissue type, the method comprising:

positioning an energy delivery portion of a catheter within the lumen adjacent the body tissue region to be heated, wherein the energy delivery portion comprises a radially expandable structure having a plurality of electrodes thereon;

expanding the radially expandable structure within the body lumen so as to radially engage the body lumen with the plurality of electrodes, wherein the plurality of electrodes engaged against the body lumen complete a plurality of circuits, wherein at least some of the circuits include a first tissue type and a second tissue type while other circuits include the first tissue type without the second tissue type;

determining a pulse characteristic in response to a thermal property of the first tissue type;

applying pulsed electrical energy with the pulse characteristic from the energy delivery portion through the plurality of circuits without identifying which of the plurality of circuits include the second tissue type so that both the first tissue type and the second tissue type are heated by thermal conduction, and while the pulsed electrical energy is applied to the other circuits at a pulsed rate that avoids thermal damage to the first tissue type, applying the same pulsed energy to the at least some circuits at the pulsed rate so as to therapeutically treat the second tissue type with heat; and

controlling the pulse characteristic in response to differing values of the thermal property of the first tissue type during application of pulsed electrical energy.

2. The method of claim 1, wherein the pulsed energy is delivered at an average rate of 0.25 to 5 watts to the body tissue region.

3. The method of claim 1, wherein each pulse of the pulsed energy provides between 4 to 45 Joules to the body tissue region.

4. The method of claim 1, wherein a time period between pulses allows the first tissue type to dissipate heat adequately to avoid thermal damage to the first tissue type by the pulsed energy.

5. The method of claim 1, wherein a time period between pulses of energy is between 0.1 to 180 seconds.

6. A catheter system for heating a body tissue region adjacent a body lumen, the region having both a first tissue and a second tissue type, the system comprising:

an elongate flexible catheter body having a proximal end and a distal end with an axis therebetween, the catheter body having a radially expandable structure near the distal end;

an energy delivery portion proximate the distal end, wherein the energy delivery portion comprises a plurality of electrodes positioned about the radially expandable structure such that when expanded within the body lumen the plurality of electrodes radially engage the body lumen, wherein the plurality of electrodes engaged against the body lumen complete a plurality of circuits, wherein at least some of the circuits include a first tissue type and a second tissue type while other circuits include the first tissue type without the second tissue type;

an energy source coupled to the energy delivery portion; and

a processor coupled to the energy source, the processor configured to control a pulse characteristic of pulsed energy transmitted from the energy source to each of the plurality of circuits without identifying which of the plurality of circuits include the second tissue type so that both the first tissue type and the second tissue type are heated by thermal conduction, wherein the energy delivery portion is configured to apply the pulsed electrical energy to the other circuits at a pulsed rate that avoids thermal damage to the first tissue type, and to apply the same pulsed energy to the at least some circuits at the pulsed rate so as to therapeutically treat the second tissue type with heat.

7. The system of claim 6, wherein the processor controls the pulsed energy such that the pulsed energy preferentially heats the second tissue type more than the first tissue type, the preferential heating induced at least in part by one or more of the following group of characteristics of the second tissue type;

lower thermal conduction than a thermal conduction of the first tissue type, lower specific heat capacity than a specific heat capacity of the first tissue type, less innate blood perfusion than an innate blood perfusion of the first tissue type, and/or larger distance away from well-perfused areas than a distance between well-perfused areas and the first tissue type.

8. The system of claim 7, wherein at least some of the group of characteristics of the second tissue type cause the first tissue type to avoid significant thermal damage, while allowing heat to build in the second tissue type.

9. The system of claim 6, wherein the pulsed energy is delivered at an average rate of 0.25 to 5 watts to the body tissue region.

10. The system of claim 6, wherein each pulse of the pulsed energy provides between 4 to 45 Joules to the body tissue region.
11. The system of claim 6, wherein an average rate of pulsed energy delivered to the body tissue region is between about 0.1 to 10.0 times the rate of energy dissipation by the first tissue type.
12. The system of claim 6, wherein a time period between pulses allows the first tissue type to dissipate its heat adequately to avoid thermal damage to the first tissue type by the pulsed energy.
13. The system of claim 6, wherein a time period between pulses of pulsed energy is between 0.1 to 180 seconds.
14. The system of claim 6, wherein a thermal time constant for the first tissue type is approximately how long a discrete volume of that given tissue will take to lose 63% of the given tissues heat while undergoing an exponential decay.
15. The system of claim 12, wherein the energy source comprises a radio frequency (RF) energy source.
16. The system of claim 6, wherein the energy source comprises a laser energy source.
17. The system of claim 16, wherein the energy delivery portion comprises at least one radially oriented window coupled to at least one optical conduit extending between the proximal end of the catheter body and the at least one window for transmission of pulsed laser energy to the body tissue region.
18. The system of claim 6, wherein the energy source comprises an ultrasound transmitter configured to deliver pulsed ultrasound energy to the body tissue region.
19. The system of claim 6, wherein the energy source comprises a microwave energy source including at least one microwave antenna configured to deliver pulsed microwave energy to the body tissue region.

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Patent Attorneys for the Applicant/Nominated Person
SPRUSON & FERGUSON

1 / 4

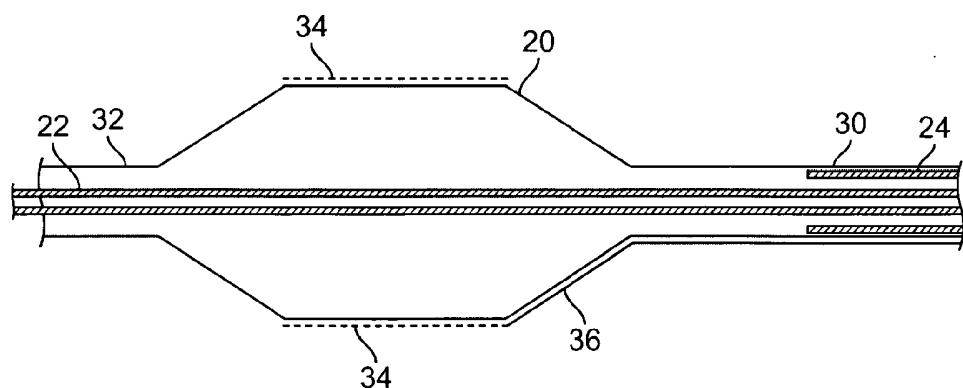
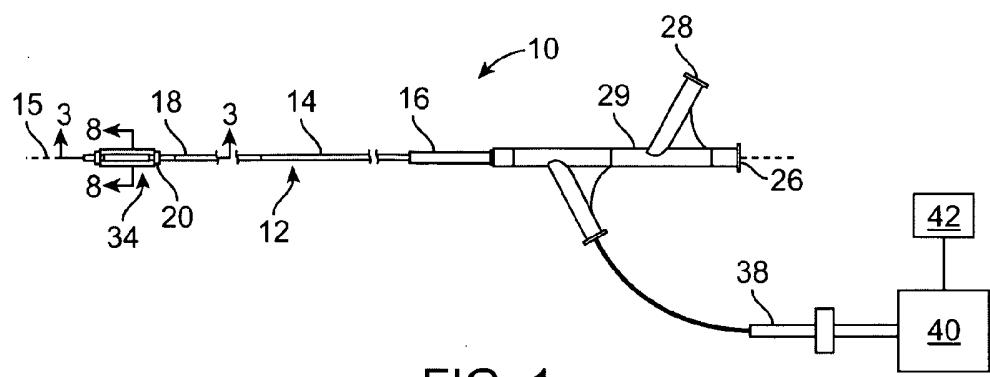


FIG. 2.

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2 / 4

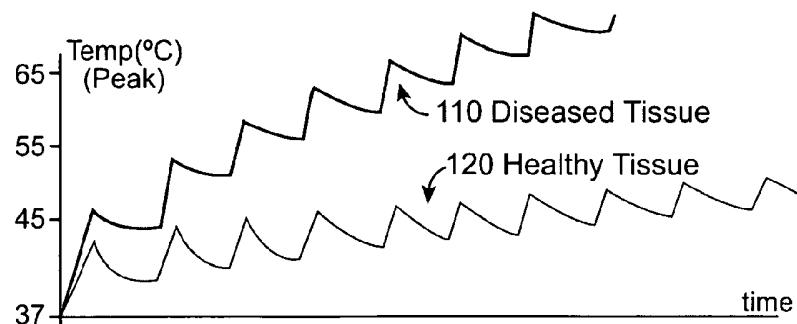


FIG. 3

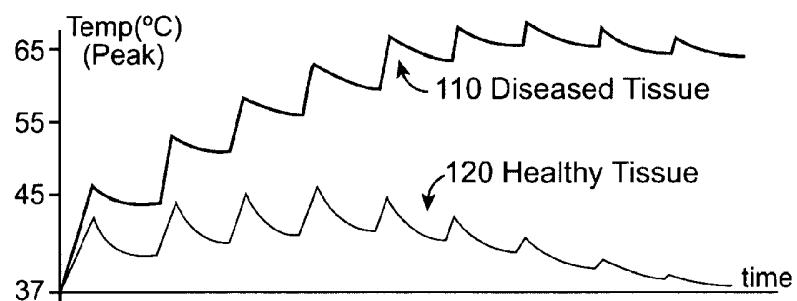
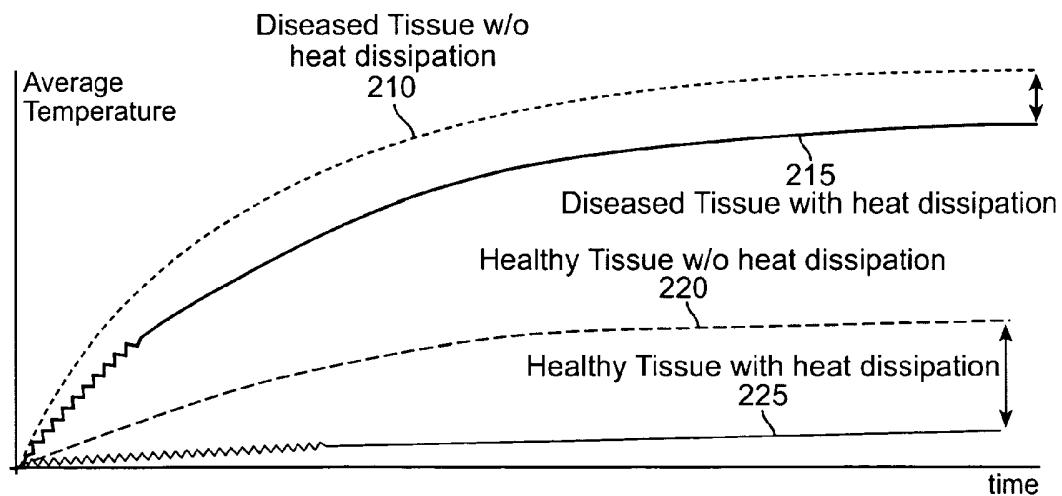


FIG. 4

FIG. 5
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3 / 4

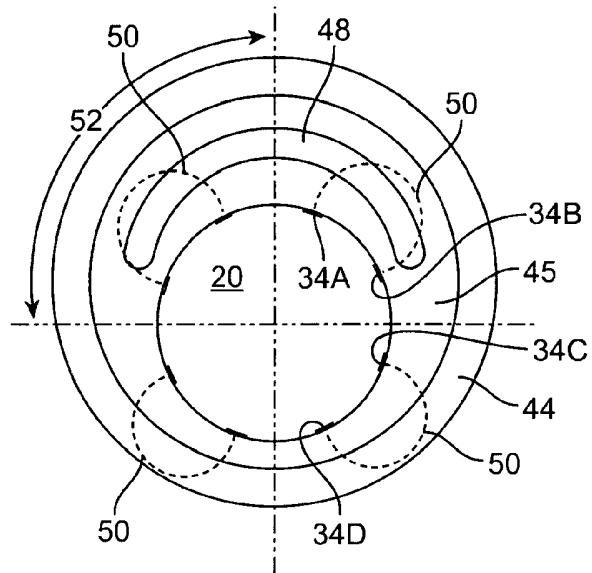


FIG. 6

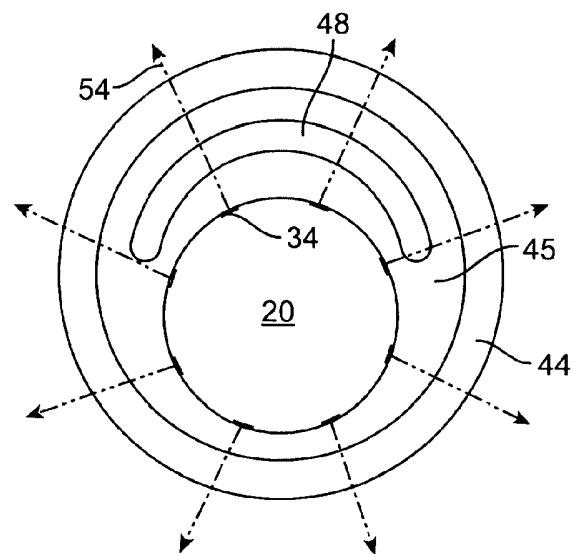


FIG. 7

4 / 4

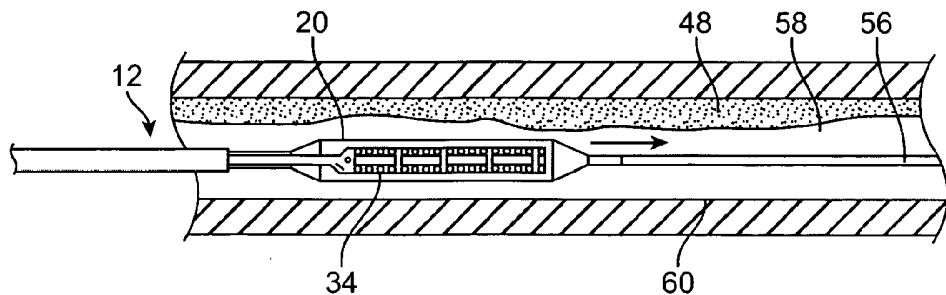


FIG. 8A

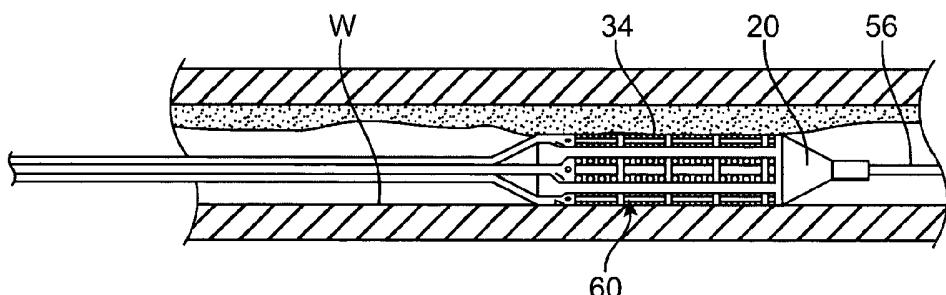


FIG. 8B

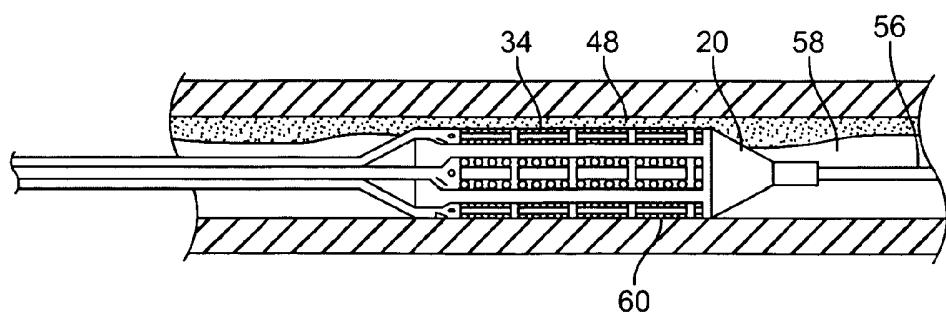


FIG. 8C