



- (51) International Patent Classification:
A61B 18/02 (2006.01)
- (21) International Application Number:
PCT/US2015/017585
- (22) International Filing Date:
25 February 2015 (25.02.2015)
- (25) Filing Language: English
- (26) Publication Language: English
- (30) Priority Data:
61/977,773 10 April 2014 (10.04.2014) US
14/265,799 30 April 2014 (30.04.2014) US
- (63) Related by continuation (CON) or continuation-in-part (CIP) to earlier application:
US 14/265,799 (CON)
Filed on 30 April 2014 (30.04.2014)
- (71) Applicant: CHANNEL MEDSYSTEMS, INC. [US/US];
5858 Horton Street, Suite 200, Emeryville, California
94608 (US).
- (72) Inventors: BURNETT, Daniel R.; 588 Teresita
Boulevard, San Francisco, California 94127 (US). COTE,
Ric; 5712 Country Club Way, Oakland, California 94802

(US). MALECKI, William W.; 91 Wildwood Gardens,
Piedmont, California 94611 (US). NEIL, Brian M.; 1434
15th Ave., San Francisco, California 94122 (US). BEAU-
LIEU, David; 7321 Lynn Ave, El Cerrito, California
94530 (US). VOILES, Benjamin D.; 248 Guerrero Street,
#202, San Francisco, California 94103 (US). LOPRESTI,
Vincent; 2040 Greenwich Street, San Francisco, California
94123 (US).

(74) Agents: HAN, Johny U. et al.; LEVINE BAGADE HAN
LLP, 2400 Geng Road, Suite 120, Palo Alto, California
94303 (US).

(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, BN, 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, IR, IS, JP, KE, KG, KN, KP, KR,
KZ, LA, LC, LK, LR, LS, LU, LY, MA, MD, ME, MG,
MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM,
PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SA, SC,
SD, SE, SG, SK, SL, SM, ST, SV, SY, TH, 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: APPARATUS AND METHODS FOR REGULATING CRYOGENIC TREATMENT

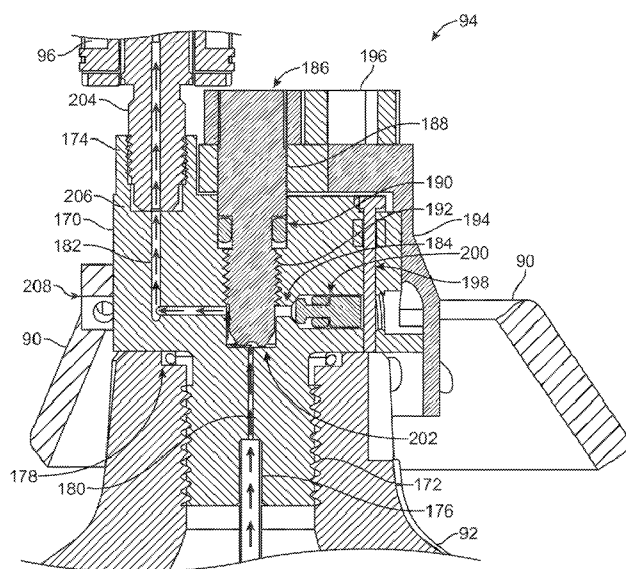


FIG. 8

(57) Abstract: Apparatus and methods for regulating
cryogenic treatments are disclosed which comprise
devices and methods for delivering controlled treatment of
a cryoablative agent. In one variation, such devices may
generally comprise an elongate probe having a distal tip
and a flexible length, at least one infusion lumen posi-
tioned through or along the elongate probe, wherein the
infusion lumen defines one or more openings along its
length, and a liner expandably enclosing the probe. An in-
flow reservoir or canister valve may be fluidly coupled
with a reservoir or canister containing the cryoablative
agent and a modulation control unit may also be fluidly
coupled with the inflow reservoir or canister valve and in
fluid communication with the at least one infusion lumen.
Additionally, a warming element may also be thermally
coupled with the reservoir or canister.



GM, KE, LR, LS, MW, MZ, NA, RW, SD, SL, ST, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, RU, TJ, TM), European (AL, 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, RS, SE, SI, SK, SM, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, KM, ML, MR, NE, SN, TD, TG).

Declarations under Rule 4.17:

- *as to applicant's entitlement to apply for and be granted a patent (Rule 4.17(ii))*
- *as to the applicant's entitlement to claim the priority of the earlier application (Rule 4.17(iii))*

Published:

- *with international search report (Art. 21(3))*

APPARATUS AND METHODS FOR REGULATING CRYOGENIC TREATMENT

CROSS-REFERENCE TO RELATED APPLICATIONS

[0001] This application claims the benefit of priority to U.S. Application No.
5 14/265,799 filed April 30, 2014 and to U.S. Provisional Application No. 61/977,773 filed
April 10, 2014, each of which is incorporated herein by reference in its entirety.

FIELD OF THE INVENTION

[0002] The present invention relates to medical devices. In particular, the present
invention relates to methods and apparatus for regulating the cryoablative treatment of
10 tissue regions.

BACKGROUND OF THE INVENTION

[0003] In the last few decades, therapeutic intervention within a body cavity or
lumen has developed rapidly with respect to delivery of energy via radiofrequency
ablation. While successful in several arenas, radiofrequency ablation has several major
15 downsides, including incomplete ablation, frequent lack of visualization during catheter
insertion, potential for overlap during treatment (with some areas receiving twice as much
energy as other areas), charring of tissues and requirements for frequent debridement,
frequent requirements for additional doses of energy after debridement, and potential
perforation of the body cavity or lumen due to the rigidity of the RF electrodes.

20 [0004] The current state of the art would benefit from minimally invasive devices
and methods which deliver thermal energy to a desired area or extract energy from a
desired area, in a consistent, controlled manner that does not char or inadvertently freeze
certain tissues or create excessive risk of unwanted organ or lumen damage.

SUMMARY OF THE INVENTION

25 [0005] Generally, devices for delivering controlled treatment may comprise an
elongate probe having a distal tip and a flexible length, at least one infusion lumen
positioned through or along the elongate probe, wherein the infusion lumen defines one or
more openings along its length, a liner expandably enclosing the probe, an inflow
reservoir or canister valve fluidly coupled with a reservoir or canister containing a

cryoablative agent, a modulation control unit fluid coupled with the inflow reservoir or canister valve and in fluid communication with the at least one infusion lumen, and a warming element thermally coupled with the reservoir or canister.

[0006] One method for utilizing the treatment assembly for cryoablatively treating tissue, e.g., uterine tissue, may generally comprising monitoring a temperature or pressure of the reservoir or canister containing a cryoablative agent, maintaining the temperature of the reservoir or canister at a predetermined level, positioning an elongate probe into a body lumen to be treated, expanding a liner enclosing the probe into contact against the body lumen, and infusing a cryoablative agent through a delivery lumen such that the cryoablative agent passes into an infusion lumen, through one or more unobstructed openings, and into contact against an interior of the liner.

[0007] In controlling or modulating the flow of the cryoablative agent, the inflow reservoir or canister valve which is fluidly coupled with the reservoir or canister may be utilized. Such a valve may generally comprising a valve body, a reservoir interface extending from the valve body and configured for fluidly coupling with the reservoir or canister containing the cryoablative agent, a modulation control interface defined along the body and configured for fluidly coupling to a modulation control interface, a valve stem seated within a valve stem channel defined within the valve body, an inflow lumen defined through the valve body and extending between the reservoir interface and the modulation control interface, where the valve stem is movable between a first position which obstructs the inflow lumen and a second position which opens the inflow lumen, a venting lumen defined through the valve body and extending between the reservoir interface and a vent opening, and a vent piston which is movable between a first position which obstructs the venting lumen and a second position which opens the venting lumen. Alternatively, the valve stem may be configured to include three positions including a first position which obstructs the inflow lumen, a second position which opens the inflow lumen, and a third optional position which opens the venting lumen.

[0008] To facilitate the liner expanding and conforming readily against the tissue walls of the uterus, the liner may be inflated with a gas or liquid. Once the elongate shaft has been introduced through the cervix and into the uterus, the distal opening of the shaft may be positioned distal to the internal os and the liner may be deployed either from within the shaft or from an external sheath. The liner may be deployed and allowed to unfurl or unwrap within the uterus. The cooling probe may be introduced through the shaft and into the liner interior. As the cryoablative agent (e.g., cryoablative fluid) is

introduced into and distributed throughout the liner interior, the exhaust catheter may also define one or more openings to allow for the cryoablative fluid to vent or exhaust from the interior of the liner.

[0009] A coolant reservoir, e.g., nitrous oxide canister, may be fluidly coupled to the handle and/or elongate shaft via a coolant valve which may be optionally controlled by the microcontroller. The coolant reservoir may be in fluid communication with the cooling probe assembly and with the interior of the balloon. Additionally, an exhaust lumen in communication with the elongate probe and having a back pressure valve may also include a pressure sensor where one or both of the back pressure sensor and/or valve may also be in communication with the microcontroller.

[0010] Yet another variation of the treatment assembly may incorporate a housing having a handle and a reservoir housing extending from and attached directly to the handle. The sheath having the liner may extend from the housing while an actuator may be located, for instance, along the handle to enable the operator to initiate the cryoablative treatment. A reservoir or canister fully containing the cryoablative fluid may be inserted and retained within the reservoir housing. The reservoir housing and/or the handle may further incorporate a reservoir engagement control which may be actuated, e.g., by rotating the control relative to the handle, to initially open fluid communication with the reservoir or canister to charge the system for treatment.

[0011] The reservoir or canister may be inserted into the reservoir housing and into secure engagement with a reservoir or canister valve which may be coupled to the reservoir engagement control. The valve may be adjusted to open the reservoir or canister for treatment or for venting of the discharged cryoablative fluid during or after treatment. An inflow modulation control unit (e.g., an actuatable solenoid mechanism) may be coupled directly to the reservoir or canister valve and the cryoablative fluid line may be coupled directly to the modulation control unit and through the sheath and into fluid communication within the liner.

[0012] The modulation control unit 96 may be in electrical communication with the microprocessor or controller via an electrical line. The inflow of the cryoablative fluid contained within the reservoir or canister may flow through an inflow line within the canister and through the reservoir or canister valve and modulation control unit and into cryoablative fluid line for introduction within the liner via infusion line. One or more pressure measurement lines which are in fluid communication with the interior of the liner may extend through the sheath and in communication with corresponding pressure

sensors which in turn are in electrical communication with microprocessor or controller via electrical lines. The pressure sensed by the measurement lines may be due (at least in part) to the expansion of the cryoablative fluid (e.g., nitrous oxide) which contacts the interior walls of the liner. Hence, microprocessor or controller may actively control the modulation control unit in a corresponding manner based on the detected pressure values within the liner sensed via pressure sensors.

[0013] To maximize patient comfort, the liner may be initially inflated with air to about, e.g., 140 mmHg, prior to the infusion of the cryoablative fluid. However, once the cryoablative fluid is introduced into the liner, the transition from air to the cryoablative fluid may create a brief fluctuation in the intracavitary pressure, e.g., spike or dip in the pressure. For instance, the pressure with which the cryoablative fluid is introduced may initially be relatively higher, e.g., about 140 mmHg. Over the course of the treatment procedure, e.g., 150 second, the pressure within the liner may result in a relatively lower pressure, e.g., about 95 mmHg.

[0014] Hence, the internal pressure within the liner during treatment may be controlled by the microprocessor or controller which may modulate the reservoir or canister valve via the modulation control unit (e.g., a solenoid valve or other mechanism) in response to the intracavitary pressures sensed by the pressure sensors. This closed-loop system may incorporate, for instance, dual pressure measuring tubes and corresponding sensors as both a redundant safety system and to also identify possible erroneous data points. The closed-loop control system can be controlled by a PID or non-PID software algorithm via the microprocessor or controller. Additionally, the modulation control unit may be used controlled by the microprocessor or controller to control the flow rate of the cryoablative fluid during the treatment procedure to optimize ablation depth and minimize the amount of cryoablative fluid needed.

[0015] During or after the treatment procedure, the discharged cryoablative fluid evacuated from the interior of the liner passes through the exhaust line which may run through the handle and reservoir housing. Hence, a system for ensuring that the discharged cryoablative fluid passing through the exhaust line is fully evaporated can be incorporated into the treatment assembly. A liquid exhaust trap which also functions as a heatsink for converting any present liquid cryogen into a gas may be integrated, for instance, directly into the reservoir housing or handle. Because the liquid exhaust trap functions as a heatsink, the trap may be fabricated from a thermally conductive material which also has a relatively large heat capacity, e.g., aluminum, copper, or other metals.

In other variations, plastics such as polycarbonate (which generally have heat capacities greater than metals such as aluminum but relatively lower thermal conductivity values) may also be utilized for fabricating the liquid exhaust trap. During use, as the discharged cryoablative fluid from the liner passes through the exhaust line and into the fluid trap
5 portion of liquid exhaust trap, any liquid form of the cryoablative fluid may collect within the fluid trap while the gaseous form may continue to be vented through the exhaust lumen and out through the evacuating exhaust line. The captured liquid may be subsequently warmed enough by contact with the liquid exhaust trap to turn into a gaseous form for venting through the exhaust line.

10 [0016] With the discharged cryoablative fluid in a completely gaseous state, the evacuating exhaust line may be vented to the surrounding environment or optionally coupled to a scavenging system to collect the discharged gas to limit exposure. Such scavenging collection systems may incorporate features such as orifices or valves to prevent any vacuum applied by the scavenging unit from interfering with the
15 backpressure within the treatment device.

[0017] In further controlling the flow of the cryoablative fluid within the treatment assembly, the reservoir or canister valve which is coupled directly to the reservoir or canister may also incorporate a number of flow control features. One variation of the reservoir or canister valve may include an integrated reservoir lumen
20 insert extending from the reservoir interface for direct insertion into the reservoir or canister to facilitate the transfer of the cryoablative fluid through the valve and into the treatment assembly. A reservoir seal may be incorporated to ensure a fluid tight seal between the reservoir or canister and the reservoir interface. The valve may include a valve body which defined pathways for normal fluid flow as well as a venting pathway
25 for emptying of the reservoir or canister.

[0018] The valve body may have the reservoir interface extending from the body for secure engagement with the reservoir or canister (e.g., via a threaded engagement). The valve body may further include a modulation control interface which defines an interface seal for securely coupling (e.g., via a threaded engagement) with a modulation
30 control interface extending from the inflow modulation control unit. A valve stem may be seated within a valve stem channel defined within the valve body. The valve stem may be secured to the valve body via a threaded engagement and a valve stem seal which ensures a fluid-tight connection between the two components. The valve stem may be

attached to a valve stem coupler which is connected to the reservoir engagement control via a control member.

[0019] During use, the reservoir engagement control may be rotated (e.g., about 45 degrees) about the reservoir housing and/or the handle. This in turn may rotate the control member and valve stem coupler which further rotates the valve stem relative to the valve body and opens the valve stem seal. The opened valve stem seal then enables the flow of the cryoablative fluid into the reservoir lumen insert and into the proximal inflow lumen located proximal to the valve stem, past the opened valve stem, and into the distal inflow lumen for further passage into the inflow modulation control unit.

[0020] Actuation of the reservoir engagement control, control member, and/or valve stem coupler may optionally send an electrical signal to the microprocessor or controller that the treatment assembly is charged with the cryoablative fluid and ready for a treatment procedure. Once the treatment procedure is completed and the inflow modulation control unit has been optionally closed to any further inflow of the cryoablative fluid, a vent pin may be actuated or pulled relative to the valve body to release a vent piston. With the vent pin secured in the valve body, the vent piston may seal a venting lumen but with the vent pin removed, the vent piston may freely translate relative to the valve body thus allowing any remaining cryoablative fluid within the reservoir or canister to vent through the venting lumen (with the valve stem still in its open position) and into the environment or into a collection reservoir, as described herein.

[0021] Yet another feature which may be optionally incorporated into the treatment assembly for controlling or modulating the flow of the cryoablative fluid may include an actuatable dome-shaped valve located within the exhaust block at a proximal end of the sheath. Although shown and described as a dome-shaped valve, such a valve is one of a variety of pneumatic and/or electro-mechanical valves that may be used to open and close the exhaust gas pathway in the assembly described herein. The exhaust block may comprise in part a body which defines an exhaust lumen in fluid communication with the interior of the liner. The exhaust lumen may also be in fluid communication with a pump/vacuum lumen which provides a channel for air for the initial inflation of the liner against the tissue surface prior to infusion of the cryoablative fluid.

[0022] The valve may generally comprise a dome-shaped flexible member attached at its periphery to the body via attachment. The flexible member may further include a seal which extends from a central portion of the concave surface of the flexible member. The flexible member may be located within a pressurization chamber which

normally exerts a pressure which is less than a deflection force required to collapse the flexible member. When the treatment assembly is used to initially puff the liner with air to force the liner into contact with the surrounding tissue, the air may pass through the pump/vacuum lumen and into the interior of the liner. The air within the pressurization chamber may also be pressurized by the same pump such that the pressure increase collapses the flexible member and forces the seal into contact against a corresponding sealing lip located at an opening of the exhaust lumen adjacent to the seal.

[0023] When the initial pressurization of the liner has been completed, the air may bleed out of the lumens as well out of the pressurization chamber allowing the flexible member to reconfigure into its opened domed shape and to release the seal from the sealing lip. This may then allow for the exhaust from the liner interior to flow through the exhaust lumen, through an exhaust chamber, and further into an exhaust lumen for venting from the treatment assembly.

[0024] In further facilitating a treatment procedure, the liner may also be configured to aid in its removal from the underlying tissue after a cryoablation treatment. After the tissue has been treated, the liner may remain frozen on the underlying uterine tissue preventing removal of the liner from the patient's body for up to several minutes. The liner may be left in the patient for a period of time after the cryoablation treatment until the tissue thaws as pulling the liner from the tissue prematurely may tear the liner; however, leaving the liner in place may increase patient discomfort. Hence, to facilitate removal of the liner from the underlying frozen tissue, a number of different warming techniques may be optionally implemented.

[0025] Circulating a warm or room temperature fluid within the liner is one method for thawing the liner and adjacent tissue to expedite the removal of the liner. A gas (e.g., air, expanded helium, etc.) may be used instead of a liquid as a warming gas may prevent the creation of a solid which could potentially block the exhaust gas pathway. Additionally, use of a warming gas may also slow the boiling-off of any remaining cryoablative fluid as boiling-off the cryoablative fluid too quickly could create a pressure spike within the liner. A liquid with a freezing point lower than the boiling point of the cryoablative fluid, such as nitrous oxide, may be utilized. It may also be possible to use a liquid which has a relatively higher boiling point than the cryoablative fluid provided that all of the cryoablative fluid has previously boiled-off. Closing the actuatable valve within the exhaust block and measuring the pressure inside the liner is one way to detect if any of the cryoablative fluid remains where an increase in pressure

would indicate the presence of liquid cryoablative fluid still boiling-off. Aside from forming an integrated fluid lumen into the liner, other mechanisms may instead be utilized to facilitate liner removal from the contacted tissue.

[0026] In these examples and any of the variations herein, a wire or heating element which may be warmed or energized (e.g., infrared) may be located on the probe shaft or positioned within the liner interior. Once the treatment procedure has been completed, the wire or heating element may be activated to warm the liner and the adjacent contacted tissue to facilitate the thawing of the tissue for removal of the liner.

[0027] Additionally and/or alternatively, the liner may be comprised of a lubricious liner or a separate non-stick coating may be applied to the liner exterior. It is not uncommon for polymers such as urethanes, especially thin films, to stick together if tightly-packed during sterilization, transportation and storage. The liner, being a thin polyurethane film compressed into a sheath, may employ a lubricious material or surface to ensure that the liner fully deploys and inflates following unsheathing.

[0028] Because optimal ablation coverage and depths may not be uniform over the entire contacted tissue region, the liner may be adjusted in thickness over particular regions of the liner to insulate predetermined tissue regions to result in tailored ablations. Ablation depths may be shallower where the liner is relatively thicker due to less efficient thermal transfer across the thicker areas.

[0029] In yet another variation, the liner may be designed with one or more predetermined weak points. If excessive tension were applied to the liner while it is frozen to tissue, the liner could tear. By locating one or more weakened regions of the liner near, e.g., the proximal connection to the probe shaft, the liner may be designed to tear specifically at the designated weakened regions which may make retrieval of the detached liner as a single piece relatively easier once the tissue fully thaws. In yet another variation of the liner, the liner may be separated into several individual liners in a multi-liner assembly.

[0030] In any of the variations described herein, the cooling probe may optionally include a compressible tip having a collapsible opening defined through the tip. The compressible tip may be positioned upon the distal tip of the probe located within the interior of the liner. Because the probe may be translatable within the liner and relative to the sheath, the tip may present a soft and atraumatic surface in the event the probe is advanced into contact against the interior of the liner and underlying tissue surface to prevent liner tears or trauma to the uterine tissue.

[0031] During a cryotherapy treatment, it is desirable to control the amount of the cryoablative fluid delivered into and through the liner. A few of the parameters which may affect the flow rate and volume of the cryoablative fluid discharged from the reservoir or canister may include temperature of the treatment assembly and reservoir or canister as well as ambient temperature in which the assembly is used as such temperatures can affect the internal pressure of the reservoir or canister. One method for controlling the starting cryogen pressure is by designing the system to operate at the high end of the temperature range and heating the reservoir or canister to a specified temperature and corresponding internal pressure. The heat could be supplied by a various mechanisms such as an electrical heating element wrapped around the reservoir or canister. In one variation, the electrical power for the heating element could be provided by a battery within the device itself.

[0032] In another variation, the electrical power may be provided by a heating cradle prior to device use. A separate warming cradle may define a receiving channel sized to receive the reservoir housing of the treatment assembly. The cradle may further include an electrical connector connected to an optional power supply (DC) (which may be recharged) and/or the cradle may be electrically connected to a stationary power supply via a power supply (AC) line. The cradle may also incorporate an optional stabilizing weight to provide for stability when the treatment assembly is docked within the receiving channel.

[0033] The treatment assembly itself may incorporate a heating element (e.g., a resistive heating element) which may be wrapped partially or entirely around the reservoir or canister. A layer of insulation may also be provided around the reservoir or canister to provide for a thermally stabilized warming environment. A temperature sensor (e.g., thermocouple, thermistor, etc.) may also be incorporated for thermal contact with the reservoir or canister for sensing the canister temperature. An electrical connector may be located correspondingly along the reservoir housing for electrically contacting the electrical connector positioned upon the cradle such that the cradle may provide electrical power to the treatment assembly when docked within the cradle receiving channel.

[0034] With the heating element and temperature sensor so coupled to the microprocessor or controller, the heating assembly may form a closed-loop system where the microprocessor or controller may be programmed via a software algorithm to control the electrical power supplied to the heating element depending upon the measured temperature of the temperature sensor such that the reservoir or canister is heated to a

predetermined temperature or maintained within a predetermined temperature range prior to a cryotherapy treatment. The insulation may accordingly slow the rate of cooling of the reservoir or canister and also extend the available time between the removal of the treatment assembly from the cradle and the initiation of the treatment.

5 BRIEF DESCRIPTION OF THE DRAWINGS

[0035] Fig. 1A shows a side view of an integrated treatment assembly.

[0036] Fig. 1B shows an example of the assembly advanced through the cervix and into the uterus where the sheath may be retracted via the handle assembly to deploy the balloon.

10 [0037] Fig. 1C shows a perspective view of a cryoablation assembly having a handle assembly which may integrate the electronics and pump assembly within the handle itself.

[0038] Fig. 1D shows the handle assembly in a perspective exploded view illustrating some of the components which may be integrated within the handle.

15 [0039] Fig. 1E shows an example of the system operation during a pre-treatment puff up process.

[0040] Fig. 1F shows an example of the system operation during a treatment process.

20 [0041] Fig. 1G shows an example of the system operation during a thawing and venting process.

[0042] Figs. 2A and 2B show cross-sectional side views of yet another variation of a cooling probe which utilizes a single infusion line in combination with a translatable delivery line.

25 [0043] Figs. 3A and 3B show top and perspective views of the expanded liner with four pairs of the open delivery ports exposed in apposed direction.

[0044] Figs. 4A to 4C show side and assembly views of another variation of the treatment assembly.

[0045] Fig. 5 shows a schematic illustration of a treatment assembly having a modulated inflow valve.

30 [0046] Fig. 6 shows a schematic illustration of a treatment assembly having a ventilation trap which also acts as a heat sink.

[0047] Figs. 7A and 7B show examples of collection systems which can be used to collect the discharged liquid or gas.

[0048] Fig. 8 shows a cross-sectional side view of a cylinder valve which may be integrated with the handle of the assembly.

5 [0049] Fig. 9 shows a cross-sectional side view of an actuatable exhaust valve which may be integrated with the handle of the assembly.

[0050] Fig. 10 shows a partial cross-sectional view of a liner having an integrated lumen for introducing a warming liquid or gas to facilitate liner removal.

10 [0051] Figs. 11A and 11B show an example for fabricating the liner having the integrated lumen.

[0052] Figs. 12A and 12B show respective top and side views of a liner having an integrated tether or wire for facilitating liner removal.

[0053] Figs. 13A and 13B show respective top and side views of the tether being tensioned and the release of the liner from the tissue surface.

15 [0054] Fig. 14 shows a top view of another variation of the liner having a tether or wire extending through a guide hole defined through a tab on the exterior of the liner.

[0055] Fig. 15 shows a top view of yet another variation of the liner having a tether or wire extending through a channel defined along the liner.

20 [0056] Fig. 16 shows a top view of yet another variation of the liner having a tether or wire extending through several guide holes.

[0057] Fig. 17 shows a top view of yet another variation of the liner having a tether or wire extending around a periphery of the liner.

[0058] Fig. 18 shows a top view of yet another variation of the liner having a tether or wire coupled to a distal end of the exhaust tubing.

25 [0059] Fig. 19 shows a top view of another variation of the liner having insulated regions defined along one or both sides of the liner.

[0060] Fig. 20 shows a top view of another variation of the liner having regions configured to have predetermined weakened areas.

30 [0061] Fig. 21 shows a top view of another variation of the assembly having multiple liners.

[0062] Figs. 22A and 22B show top views of another variation of the cooling probe having a compressible tip to present an atraumatic surface to the liner interior and tissue surface.

[0063] Fig. 23 shows a partial cross-sectional side view of a treatment assembly incorporating particular materials to facilitate positioning of a probe relative to the interior of a deployed liner.

[0064] Fig. 24 shows a schematic illustration of a treatment assembly optionally including a temperature controlled cradle.

DETAILED DESCRIPTION OF THE INVENTION

[0065] The cooling probe 22 as well as the balloon assembly may be variously configured, for instance, in an integrated treatment assembly 10 as shown in the side view of Fig. 1A. In this variation, the assembly 10 may integrate the elongate shaft 18 having the liner or balloon 20 extending therefrom with the cooling probe 22 positioned translatable within the shaft 18 and liner 20. A separate translatable sheath 12 may be positioned over the elongate shaft 18 and both the elongate shaft 18 and sheath 12 may be attached to a handle assembly 14. The handle assembly 14 may further comprise an actuator 16 for controlling a translation of the sheath 12 for liner 20 delivery and deployment.

[0066] With the sheath 12 positioned over the elongate shaft 18 and liner 20, the assembly 10 may be advanced through the cervix and into the uterus UT where the sheath 12 may be retracted via the handle assembly 14 to deploy the liner 20, as shown in Fig. 1B. As described above, once the liner 20 is initially deployed from the sheath 12, it may be expanded by an initial burst of a gas, e.g., air, carbon dioxide, etc., or by the cryoablative fluid. In particular, the tapered portions of the liner 20 may be expanded to ensure contact with the uterine cornu. The handle assembly 14 may also be used to actuate and control a longitudinal position of the cooling probe 22 relative to the elongate shaft 18 and liner 20 as indicated by the arrows.

[0067] In another variation of the treatment assembly, Fig. 1C shows a perspective view of a cryoablation assembly having a handle assembly 24 which may integrate the electronics and pump assembly 28 within the handle itself. An exhaust tube 26 may also be seen attached to the handle assembly 24 for evacuating exhausted or excess cryoablative fluid or gas from the liner 20. Any of the cryoablative fluids or gases described herein may be utilized, e.g., compressed liquid-to-gas phase change of a compressed gas such as nitrous oxide (N₂O), carbon dioxide (CO₂), Argon, etc. The cooling probe 22 may be seen extending from sheath 12 while surrounded or enclosed by

the liner or balloon 20. Hence, the handle assembly 24 with coupled cooling probe 22 and liner 20 may provide for a single device which may provide for pre-treatment puff-up or inflation of the liner 20, active cryoablation treatment, and/or post-treatment thaw cycles.

5 [0068] The handle assembly 24 may also optionally incorporate a display for providing any number of indicators and/or alerts to the user. For instance, an LCD display may be provided on the handle assembly 24 (or to a separate control unit connected to the handle assembly 24) where the display counts down the treatment time in seconds as the ablation is occurring. The display may also be used to provide
10 measured pressure or temperature readings as well as any number of other indicators, symbols, or text, etc., for alerts, instructions, or other indications. Moreover, the display may be configured to have multiple color-coded outputs, e.g., green, yellow, and red. When the assembly is working through the ideal use case, the LED may be displayed as a solid green color. When the device requires user input (e.g. when paused and needing the
15 user to press the button to re-start treatment) the LED may flash or display yellow. Additionally, when the device has faulted and treatment is stopped, the LED may flash or display a solid red color.

[0069] Fig. 1D shows the handle assembly 24 in a perspective exploded view to illustrate some of the components which may be integrated within the handle 24. As
20 shown, the liner 20 and sheath 12 may be coupled to a sheath bearing assembly 32 and slider base block assembly 34 for controlling the amount of exposed treatment length along the cooling probe 22 (and as described in further detail below). An actuatable sheath control 36 may be attached to the slider base block assembly 34 for manually controlling the treatment length of the cooling probe 22 as well. Along with the
25 electronics and pump assembly 28 (which may optionally incorporate a programmable processor or controller in electrical communication with any of the mechanisms within the handle 24), an exhaust valve 30 (e.g., actuated via a solenoid) may be coupled to the exhaust line 26 for controlling not only the outflow of the exhausted cryoablation fluid or gas but also for creating or increasing a backpressure during treatment, as described in
30 further detail below.

[0070] In one example of how the handle assembly 24 may provide for treatment, Figs. 1E to 1G illustrate schematic side views of how the components may be integrated and utilized with one another. As described herein, once the sheath 12 and/or liner 20 has been advanced and initially introduced into the uterus, the liner 20 may be expanded or

inflated in a pre-treatment puff up to expand the liner 20 into contact against the uterine tissue surfaces in preparation for a cryoablation treatment. As illustrated in the side view of Fig. 1E, a pump 38 integrated within the handle assembly 24 may be actuated and a valve 42 (e.g., actuatable or passive) fluidly coupled to the pump 38 may be opened (as indicated schematically by an "O" over both the pump 38 and valve 42) such that ambient air may be drawn in through, e.g., an air filter 40 integrated along the handle 24, and passed through an air line 44 within the handle and to an exhaust block 46. The exhaust block 46 and air line 44 may be fluidly coupled to the tubular exhaust channel which extends from the handle 24 which is further attached to the cooling probe 22. As the air is introduced into the interior of the liner 20 (indicated by the arrows), the liner 20 may be expanded into contact against the surrounding uterine tissue surface.

[0071] A cryoablative fluid line 48 also extending into and integrated within the handle assembly 24 may be fluidly coupled to an actuatable valve 50, e.g., actuated via a solenoid, which may be manually closed or automatically closed (as indicated schematically by an "X" over the valve 50) by a controller to prevent the introduction of the cryoablative fluid or gas into the liner 20 during the pre-treatment liner expansion. An infusion line 52 may be fluidly coupled to the valve 50 and may also be coupled along the length of the sheath 12 and probe 22, as described in further detail below. The exhaust valve 30 coupled to the exhaust line 26 may also be closed (as indicated schematically by an "X" over the valve 30) manually or automatically by the controller to prevent the escape of the air from the exhaust block 46.

[0072] During this initial liner expansion, the liner 20 may be expanded in a gradual and controlled manner to minimize any pain which may be experienced by the patient in opening the uterine cavity. Hence, the liner 20 may be expanded gradually by metering in small amounts of air. Optionally, the pump 38 may be programmed and controlled by a processor or microcontroller to expand the liner 20 according to an algorithm (e.g., e.g. ramp-up pressure quickly to 10 mm Hg and then slow-down the ramp-up as the pressure increases to 85 mm Hg) which may be stopped or paused by the user. Moreover, the liner 20 may be expanded to a volume which is just sufficient to take up space within the uterine cavity. After the initial increase in pressure, the pressure within the liner 20 may be optionally increased in bursts or pulses. Moreover, visualization (e.g., via a hysteroscope or abdominal ultrasound) may be optionally used during the controlled gradual expansion to determine when the uterine cavity is fully open and requires no further pressurization. In yet another variation, the liner 20 may be

cyclically inflated and deflated to fully expand the liner. The inflations and deflations may be partial or full depending upon the desired expansion.

[0073] In yet another alternative variation, the system could also use an amount of air pumped into the liner 20 as a mechanism for detecting whether the device is in a false passage of the body rather than the uterine cavity to be treated. The system could use the amount of time that the pump 38 is on to track how much air has been pushed into the liner 20. If the pump 38 fails to reach certain pressure levels within a predetermined period of time, then the controller may indicate that the device is positioned within a false passage. There could also be a limit to the amount of air allowed to be pushed into the liner 20 as a way to detect whether the probe 22 has been pushed, e.g., out into the peritoneal cavity. If too much air is pushed into the liner 20 (e.g., the volume of air tracked by the controller exceeds a predetermined level) before reaching certain pressures, then the controller may indicate the presence of a leak or that the liner 20 is not fully constrained by the uterine cavity. The liner 20 may also incorporate a release feature which is configured to rupture if the liner 20 is not constrained such that if the system attempts to pump up the liner 20 to treatment pressure (e.g., 140 mmHg), the release feature will rupture before reaching that pressure.

[0074] Once the liner 20 has been expanded sufficiently into contact against the uterine tissue surface, the cryoablation treatment may be initiated. As shown in the side view of Fig. 1F, the air pump 38 may be turned off and the valve 42 may be closed (as indicated schematically by an "X" over the pump 38 and valve 42) to prevent any further infusion of air into the liner 20. With the cryoablative fluid or gas pressurized within the line 48, valve 50 may be opened (as indicated schematically by an "O" over the valve 50) to allow for the flow of the cryoablative fluid or gas to flow through the infusion line 52 coupled to the valve 50. Infusion line 52 may be routed through or along the sheath 12 and along the probe 22 where it may introduce the cryoablative fluid or gas within the interior of liner 20 for infusion against the liner 20 contacted against the surrounding tissue surface.

[0075] During treatment or afterwards, the exhaust valve 30 may also be opened (as indicated schematically by an "O" over the valve 30) to allow for the discharged fluid or gas to exit or be drawn from the liner interior and proximally through the cooling probe 22, such as through the distal tip opening. The fluid or gas may exit from the liner 20 due to a pressure differential between the liner interior and the exhaust exit and/or the fluid or gas may be actively drawn out from the liner interior, as described in further detail herein.

The spent fluid or gas may then be withdrawn proximally through the probe **22** and through the lumen surrounded by the sheath **12**, exhaust block **46**, and the exhaust tube **26** where the spent fluid or gas may be vented. With the treatment fluid or gas thus introduced through infusion line **52** within the liner **20** and then withdrawn, the cryoablative treatment may be applied uninterrupted.

[0076] Once a treatment has been completed, the tissue of the uterine cavity may be permitted to thaw. During this process, the cryoablative fluid delivery is halted through the infusion line **52** by closing the valve **50** (as indicated schematically by an "X" over the valve **50**) while continuing to exhaust for any remaining cryoablative fluid or gas remaining within the liner **20** through probe **22**, through the lumen surrounded by sheath **12**, and exhaust line **26**, as shown in Fig. 1G. Optionally, the pump **38** and valve **42** may be cycled on and off and the exhaust valve **30** may also be cycled on and off to push ambient air into the liner **20** to facilitate the thawing of the liner **20** to the uterine cavity. Optionally, warmed or room temperature air or fluid (e.g., saline) may also be pumped into the liner **20** to further facilitate thawing of the tissue region.

[0077] As the spent cryoablative fluid or gas is removed from the liner **20**, a drip prevention system may be optionally incorporated into the handle. For instance, a passive system incorporating a vented trap may be integrated into the handle which allows exhaust gas to escape but captures any vented liquid. The exhaust line **26** may be elongated to allow for any vented liquid to evaporate or the exhaust line **26** may be convoluted to increase the surface area of the exhaust gas tube to promote evaporation.

[0078] Alternatively, an active system may be integrated into the handle or coupled to the handle **24** where a heat sink may be connected to a temperature sensor and electrical circuit which is controlled by a processor or microcontroller. The heat sink may promote heat transfer and causes any liquid exhaust to evaporate. When the temperature of the heat sink reaches the boiling temperature of, e.g., nitrous oxide (around -86 °C), the handle may be configured to slow or stop the delivery of the cryoablative fluid or gas to the uterine cavity.

[0079] The pre-treatment infusion of air as well as the methods for treatment and thawing may be utilized with any of the liner, probe, or apparatus variations described herein. Moreover, the pre-treatment, treatment, or post-treatment procedures may be utilized altogether in a single procedure or different aspects of such procedures may be used in varying combinations depending upon the desired results.

[0080] Additionally and/or optionally, the handle 24 may incorporate an orientation sensor to facilitate maintaining the handle 24 in a desirable orientation for treatment. One variation may incorporate a ball having a specific weight covering the exhaust line 26 such that when the handle 24 is held in the desirable upright orientation, the treatment may proceed uninterrupted. However, if the handle 24 moved out of its desired orientation, the ball may be configured to roll out of position and trigger a visual and/or auditory alarm to alert the user. In another variation, an electronic gyroscopic sensor may be used to maintain the handle 24 in the desired orientation for treatment.

[0081] Figs. 2A and 2B show cross-sectional side views of yet another variation of a cooling probe which utilizes a single infusion line in combination with a translatable delivery line. To accommodate various sizes and shapes of uterine cavities, the cooling probe may have a sliding adjustment that may be set, e.g., according to the measured length of the patient's uterine cavity. The adjustment may move along the sheath along the exhaust tube as well as the delivery line within the infusion line. The sheath may constrain the liner 20 and also control its deployment within the cavity.

[0082] In this variation, an infusion line 52 (as described above) may pass from the handle assembly and along or within the sheath and into the interior of liner 20. The infusion line 52 may be aligned along the probe 22 such that the infusion line 52 is parallel with a longitudinal axis of the probe 22 and extends towards the distal tip 66 of the probe 22. Moreover, the infusion line 52 may be positioned along the probe 22 such that the line 52 remains exposed to the corners of the liner 20 which extend towards the cornua. With the infusion line 52 positioned accordingly, the length of the line 52 within the liner 20 may have multiple openings formed along its length which act as delivery ports for the infused cryoablative fluid or gas. A separate translating delivery line 64, e.g., formed of a Nitinol tube defining an infusion lumen therethrough, may be slidably positioned through the length of the infusion line 52 such that the delivery line 64 may be moved (as indicated by the arrows in Fig. 2A) relative to the infusion line 52 which remains stationary relative to the probe 22.

[0083] The openings along the length of the infusion line 52 may be positioned such that the openings are exposed to the sides of the interior of the liner 20, e.g., cross-drilled. As the cryoablative fluid or gas is introduced through the delivery line 64, the infused cryoablative fluid or gas 68 may pass through the infusion line 52 and then out through the openings defined along the infusion line 52. By adjusting the translational position of the delivery line 64, the delivery line 64 may also cover a selected number of

the openings resulting in a number of open delivery ports **60** as well as closed delivery ports **62** which are obstructed by the delivery line **64** position relative to the infusion line **52**, as shown in the top view of Fig. 2B.

[0084] By translating the delivery line **64** accordingly, the number of open delivery ports **60** and closed delivery ports **62** may be adjusted depending on the desired treatment length and further ensures that only desired regions of the uterine tissue are exposed to the infused cryoablative fluid or gas **68**. Once the number of open delivery ports **60** has been suitably selected, the infused cryoablative fluid or gas **68** may bypass the closed delivery ports **62** obstructed by the delivery line **64** and the fluid or gas may then be forced out through the open delivery ports **60** in a transverse direction as indicated by the infusion spray direction **70**. The terminal end of the infusion line **52** may be obstructed to prevent the distal release of the infused fluid or gas **68** from its distal end. Although in other variations, the terminal end of the infusion line **52** may be left unobstructed and opened.

[0085] Figs. 3A and 3B show top and perspective views of the expanded liner **20** with four pairs of the open delivery ports **60** exposed in apposed direction. Because the infused fluid or gas **68** may be injected into the liner **20**, e.g., as a liquid, under relatively high pressure, the injected cryoablative liquid may be sprayed through the open delivery ports **60** in a transverse or perpendicular direction relative to the cooling probe **22**. The laterally infused cryoablative fluid **70** may spray against the interior of the liner **20** (which is contacted against the surrounding tissue surface) such that the cryoablative liquid **70** coats the interior walls of the liner **20** due to turbulent flow causing heavy mixing. As the cryoablative liquid **70** coats the liner surface, the sprayed liquid **70** may absorb heat from the tissue walls causing rapid cooling of the tissue while also evaporating the liquid cryogen to a gas form that flows out through the cooling probe **22**. This rapid cooling and evaporation of the cryoablative liquid **70** facilitates the creation of a fast and deep ablation over the tissue. During treatment, the temperature within the cavity typically drops, e.g., -86° C, within 2-3 seconds after the procedure has started. While the interior walls of the liner **20** are first coated with the cryoablative liquid **70**, a portion of the cryoablative liquid **70** may no longer change phase as the procedure progresses.

[0086] While four pairs of the open delivery ports **60** are shown, the number of exposed openings may be adjusted to fewer than four pairs or more than four pairs depending on the positioning of the delivery line **64** and also the number of openings defined along the infusion line **52** as well as the spacing between the openings.

Moreover, the positioning of the openings may also be adjusted such that the sprayed liquid **70** may spray in alternative directions rather than laterally as shown. Additionally and/or alternatively, additional openings may be defined along other regions of the infusion line **52**.

5 **[0087]** Further variations of the treatment assembly features and methods which may be utilized in combination with any of the features and methods described herein may be found in the following patent applications:

US Pat. App. 13/361,779 filed January 30, 2012 (US Pub. 2012/0197245);
US Pat. App. 13/900,916 filed May 23, 2013 (US Pub. 2013-0296837);
10 US Pat. App. 14/019,898 filed September 6, 2013 (US Pub. 2014/0012156);
US Pat. App. 14/019,928 filed September 6, 2013 (US Pub. 2014/005648);
US Pat. App. 14/020,265 filed September 6, 2013 (US Pub. 2014/0005649);
US Pat. App. 14/020,306 filed September 6, 2013 (US Pub. 2014/0025055);
US Pat. App. 14/020,350 filed September 6, 2013 (US Pub. 2014/0012244);
15 US Pat. App. 14/020,397 filed September 6, 2013 (US Pub. 2014/0012243);
US Pat. App. 14/020,452 filed September 6, 2013 (US Pub. 2014/0005650);
US Pat. App. 14/086,050 filed November 21, 2013 (US Pub. 2014/0074081);
US Pat. App. 14/086,088 filed November 21, 2013 (US Pub. 2014/0088579); and
US Pat. App. 14/029,641 filed September 17, 2013.

20 **[0088]** Each of the patent applications above is incorporated herein by reference in its entirety and for any purpose herein.

[0089] Yet another variation of the treatment assembly **80** is shown in the side and partial cross-sectional side views of Figs. 4A and 4B which illustrate a housing **82** having a handle **84** and a reservoir housing **88** extending from and attached directly to the
25 handle **84**. Fig. 4C further illustrates a perspective assembly view of the treatment assembly **80** and some of its components contained internally.

[0090] The sheath **12** having the liner **20** may extend from the housing **82** while an actuator **86** may be located, for instance, along the handle **84** to enable the operator to initiate the cryoablative treatment. A reservoir or canister **92** fully containing the
30 cryoablative agent (as described herein) may be inserted and retained within the reservoir housing **88**. The reservoir housing **88** and/or the handle **84** may further incorporate a reservoir engagement control **90** which may be actuated, e.g., by rotating the control **90** relative to the handle **84**, to initially open fluid communication with the reservoir or canister **92** to charge the system for treatment.

[0091] The reservoir or canister 92 may be inserted into the reservoir housing 88 and into secure engagement with a reservoir or canister valve 94 which may be coupled to the reservoir engagement control 90. The valve 94 may be adjusted to open the reservoir or canister 92 for treatment or for venting of the discharged cryoablative agent during or after treatment. An inflow modulation control unit 96 (e.g., an actuatable solenoid mechanism) may be coupled directly to the reservoir or canister valve 94 and the cryoablative fluid line 48 may be coupled directly to the modulation control unit 96 and through the sheath 12 and into fluid communication within the liner 20, as described herein.

[0092] During or after treatment, the discharged cryoablative fluid may be evacuated through the exhaust block 46 contained within the housing and then through the exhaust line 98 coupled to the exhaust block 46. The exhaust line 98 may extend through the handle 84 and the reservoir housing 88 and terminate at an exhaust line opening 100 which may be attached to another exhaust collection line, as further described herein.

[0093] INFLOW MODULATION CONTROL

[0094] The reservoir or canister valve 94 and modulation control unit 96 are further illustrated in the schematic view of Fig. 5 which shows the modulation control unit 96 in electrical communication with microprocessor or controller 110 (which may be part of the electronics and pump assembly 28) via electrical line 116. As shown, the inflow of the cryoablative agent 120 contained within the reservoir or canister 92 may flow through an inflow line 122 within the canister 92 and through the reservoir or canister valve 94 and modulation control unit 96 and into cryoablative fluid line 48 for introduction within the liner 20 via infusion line 52, as described herein.

[0095] One or more pressure measurement lines 118 which are in fluid communication with the interior of the liner 20 may extend through the sheath 12 and in communication with corresponding pressure sensors 112 which in turn are in electrical communication with microprocessor or controller 110 via electrical lines 114. The pressure sensed by the measurement lines 118 may be due (at least in part) to the expansion of the cryoablative agent 120 (e.g., nitrous oxide) which contacts the interior walls of the liner 20, as described herein. Hence, microprocessor or controller 110 may actively control the modulation control unit 96 in a corresponding manner based on the detected pressure values within the liner 20 sensed via pressure sensors 112.

[0096] It is desirable to control the pressure within the liner **20** when positioned within the uterine cavity to minimize patient discomfort while simultaneously ensuring that the liner **20** remains fully deployed and in contact with the endometrial tissue. A pressure of about, e.g., 140 mmHg, is near the maximum pressure typically used during a hysteroscopy and is well-tolerated by most patients. However, other endometrial ablation devices may have operating pressures of up to 180 mmHg to 220 mmHg (or more) but such therapies typically require that the patients receive sedation due to hyperthermic ablation and high uterine pressures.

[0097] Hence, to maximize patient comfort, the liner **20** may be initially inflated with air to about, e.g., 140 mmHg, prior to the infusion of the cryoablative agent **120**, as also described herein. However, once the cryo cryoablative agent **120** is introduced into the liner **20**, the transition from air to the cryoablative agent **120** may create a brief fluctuation in the intracavitary pressure, e.g., spike or dip in the pressure. For instance, the pressure with which the cryoablative agent **120** is introduced may initially be relatively higher, e.g., about 150 mmHg. Over the course of the treatment procedure, e.g., 150 second, the pressure within the liner **20** may result in a relatively lower pressure, e.g., about 95 mmHg.

[0098] Various factors may have an influence on the pressure fluctuation over the course of the treatment procedure. For example, the initial pressure within the reservoir or canister **92** (e.g., nitrous oxide cylinder or tank) may have an impact on the pressure within the liner **20** during treatment. Conditions such as ambient temperature and/or temperature of the reservoir or canister may also have an influence. For instance, the warmer the device temperature, the greater the cylinder pressure and cryoablative agent **120** flow rate and, correspondingly, the greater the intrauterine pressure.

[0099] Hence, the internal pressure within the liner **20** during treatment may be controlled by the microprocessor or controller **110** which may modulate the reservoir or canister valve **94** via the modulation control unit **96** (e.g., a solenoid valve or other mechanism) in response to the intracavitary pressures sensed by the pressure sensors **112**. This closed-loop system may incorporate, for instance, dual pressure measuring tubes **118** and corresponding sensors **112** as both a redundant safety system and to also identify possible erroneous data points. The closed-loop control system can be controlled by a PID or non-PID software algorithm via the microprocessor or controller **110**. Additionally, the modulation control unit **96** may be used controlled by the microprocessor or controller **110** to control the flow rate of the cryoablative agent **120**

during the treatment procedure to optimize ablation depth and minimize the amount of cryoablative agent needed.

[0100] EVACUATION CONTROL

[0101] During or after the treatment procedure, the discharged cryoablative agent

5 **120** evacuated from the interior of the liner **20** passes through the exhaust line **98** which may run through the handle **84** and reservoir housing **88**, as further shown in the schematic illustration in Fig. 6. At some point during a cryoablation procedure, it may not be uncommon for the cryoablative agent still in a liquid phase to appear in the exhaust gas passing through exhaust line **98**. Having the exhaust remain in droplets of liquid
10 nitrous oxide could potentially pose a hazard to patients by coming into contact with the patient's or user's skin and larger drops of the liquid cryoablative agent could potentially cause burns. Hence, a system for ensuring that the discharged cryoablative agent passing through the exhaust line **98** is fully evaporated can be incorporated into the treatment assembly.

15 **[0102]** In the variation shown, a liquid exhaust trap **130** which may also function as a heatsink for converting any present liquid cryogen into a gas may be integrated, for instance, directly into the reservoir housing **88** or handle **84**. Such a liquid exhaust trap **130** may generally include a fluid trap **134** near the bottom portion of the liquid exhaust trap **130** where the exhaust line opening **100** may be positioned. An exhaust lumen **132**
20 may extend within the liquid exhaust trap **130** from the fluid trap **134** and the exhaust lumen **132** may further define an opening which is clear of any fluid which may be collected within the fluid trap **134**. An evacuating exhaust line **140** may be coupled to an opening **138** in communication with the exhaust lumen **132**.

[0103] Because the liquid exhaust trap **130** may function as a heatsink, the trap
25 **130** may be fabricated from a thermally conductive material which also has a relatively large heat capacity, e.g., aluminum, copper, or other metals. In other variations, plastics such as polycarbonate (which generally have heat capacities greater than metals such as aluminum but relatively lower thermal conductivity values) may also be utilized for fabricating the liquid exhaust trap **130**. Other factors such as weight and manufacturing
30 processes are further considerations in designing the liquid exhaust trap **130**.

Additionally, a fan may also be incorporated into the liquid exhaust trap **130** to facilitate thermal energy exchange with the environment. During use, as the discharged cryoablative agent from the liner **20** passes through the exhaust line **98** and into the fluid trap **134** portion of liquid exhaust trap **130**, any liquid **136** form of the cryoablative agent

may collect within the fluid trap **134** while the gaseous form may continue to be vented through the exhaust lumen **132** and out through the evacuating exhaust line **140**. The captured liquid **136** may be subsequently warmed enough by contact with the liquid exhaust trap **130** to turn into a gaseous form for venting through the exhaust lumen **132**.

5 **[0104]** Additionally and/or alternatively with respect to the liquid exhaust trap **130**, the evacuating exhaust line **140** may form a length of tubing, e.g., 5 ft., which may optionally be convoluted in configuration and which could be used to provide sufficient surface area and a pathway to facilitate heat exchange with the environment and promote the evaporation of any liquid cryoablative agent.

10 **[0105]** With the discharged cryoablative agent in a completely gaseous state, the evacuating exhaust line **140** may be vented to the surrounding environment or optionally coupled to a scavenging system to collect the discharged gas to limit exposure. Figs. 7A and 7B show assembly views of examples of collection bags which may be optionally used with the treatment assembly. Scavenging systems may incorporate features such as
15 orifices or valves to prevent any vacuum applied by the scavenging unit from interfering with the backpressure within the treatment device.

[0106] Fig. 7A shows an inflating collection bag **150** which is expandable in width coupled to the evacuating exhaust line **140** via a disconnect valve **152** (e.g., unidirectional valve). The collection bag **150**, which may be reusable or disposable, may
20 be supported via a pole **156** and may also incorporate a release plug **154** which may allow for the venting of the collected gas during or after a treatment procedure is completed.

[0107] Similarly, Fig. 7B shows an accordion-type collector **160** also supported via a pole **156** and a connector **166** attached to the collector **160**. The evacuating exhaust line **140** may be removably coupled to the collector **160** via a disconnect valve **162** (e.g.,
25 unidirectional valve) and may also incorporate a release plug **164** for venting any collected gas during or after a treatment procedure. The vertically-expanding collector **160** may define a hollow passageway through the center of the vertical bellows which allows for the connector **166** (e.g., rigid rod or flexible cord) to pass through and support the base of the collector **160**. The connector **166** also prevents the collector **160** from
30 falling over to a side when inflating. As the gas enters through the bottom of the collector **160**, the bellow may inflate upward.

[0108] INFLOW AND VENTING CONTROL

[0109] In further controlling the flow of the cryoablative agent within the treatment assembly, the reservoir or canister valve **94** which is coupled directly to the

reservoir or canister **92** may also incorporate a number of flow control features. Fig. 8 shows a cross-sectional side view of one variation of the reservoir or canister valve **94** which may include an integrated reservoir lumen insert **176** extending from the reservoir interface **172** for direct insertion into the reservoir or canister **92** to facilitate the transfer of the liquid cryoablative agent through the valve **94** and into the treatment assembly. A reservoir seal **178** may be incorporated to ensure a fluid tight seal between the reservoir or canister **92** and the reservoir interface **172**. The valve **94** may include a valve body **170** which defined pathways for normal fluid flow as well as a venting pathway for emptying of the reservoir or canister **92**.

[0110] The valve body **170** may have the reservoir interface **172** extending from the body **170** for secure engagement with the reservoir or canister **92** (e.g., via a threaded engagement). The valve body **170** may further include a modulation control interface **174** which defines an interface seal **206** for securely coupling (e.g., via a threaded engagement) with a modulation control coupler **204** extending from the inflow modulation control unit **96**. A valve stem **186** may be seated within a valve stem channel **188** defined within the valve body **170**. The valve stem **186** may be secured to the valve body **170** via a threaded engagement **192** and a valve stem seal **190** which ensures a fluid-tight connection between the two components. The valve stem **186** may be attached to a valve stem coupler **196** which is connected to the reservoir engagement control **90** via a control member **194**.

[0111] During use, the reservoir engagement control **90** may be rotated (e.g., about 45 degrees) about the reservoir housing **88** and/or the handle **84**. This in turn may rotate the control member **194** and valve stem coupler **196** which further rotates the valve stem **186** relative to the valve body **170** and opens the valve stem seal **202**. The opened valve stem seal **202** then enables the flow of the cryoablative agent into the reservoir lumen insert **176** and into the proximal inflow lumen **180** located proximal to the valve stem **186**, past the opened valve stem seal **202**, and into the distal inflow lumen **182** for further passage into the inflow modulation control unit **96**.

[0112] Actuation of the reservoir engagement control **90**, control member **194**, and/or valve stem coupler **196** may optionally send an electrical signal to the microprocessor or controller **110** that the treatment assembly **80** is charged with the cryoablative agent and ready for a treatment procedure. Once the treatment procedure is completed and the inflow modulation control unit **96** has been optionally closed to any further inflow of the cryoablative agent, a vent pin **198** may be actuated or pulled relative

to the valve body **170** to release a vent piston **200**. With the vent pin **198** secured in the valve body **170**, the vent piston **200** may seal a venting lumen **184** but with the vent pin **198** removed, the vent piston **200** may freely translate relative to the valve body **170** thus allowing any remaining cryoablative agent within the reservoir or canister **92** to vent
5 through the venting lumen **184** (with the valve stem **186** still in its open position) and into the environment or into a collection reservoir, as described herein.

[0113] Additionally, a pressure relief mechanism **208** may be optionally incorporated into the valve body **170** to function as a burst valve or other pressure release mechanism for safety purposes.

10 [0114] Yet another feature which may be optionally incorporated into the treatment assembly for opening and closing the exhaust gas pathway to facilitate gradual pressurization of the liner and uterine cavity with filtered air as well as the application of vacuum to the liner and uterine cavity following the infusion of the cryoablative fluid may be seen in the partial cross-sectional side view of Fig. 9 which illustrates an
15 actuatable dome-shaped valve located within the exhaust block **46** at a proximal end of the sheath **12**. The exhaust block **46** may comprise in part a body **210** which defines an exhaust lumen **212** in fluid communication with the interior of the liner **20**. The exhaust lumen **212** may also be in fluid communication with a pump/vacuum lumen **214** which provides a channel for air for the initial inflation of the liner **20** against the tissue surface
20 prior to infusion of the cryoablative agent.

[0115] Although shown and described as a dome-shaped valve, such a valve is one of a variety of pneumatic and/or electro-mechanical valves that may be used to open and close the exhaust gas pathway in the assembly described herein. The valve may generally comprise a dome-shaped flexible member **220** attached at its periphery to the
25 body **210** via attachment **226**. The flexible member **220** may further include a seal **224** which extends from a central portion of the concave surface of the flexible member **220**. The flexible member **220** may be located within a pressurization chamber **222** which normally exerts a pressure which is less than a deflection force required to collapse the flexible member **220**. When the treatment assembly is used to initially puff the liner **20**
30 with air to force the liner **20** into contact with the surrounding tissue, the air may pass through the pump/vacuum lumen **214** and into the interior of the liner **20**. The air within the pressurization chamber **222** may also be pressurized by the same pump such that the pressure increase collapses the flexible member **220** and forces the seal **224** into contact

against a corresponding sealing lip 228 located at an opening of the exhaust lumen 212 adjacent to the seal 224.

[0116] When the initial pressurization of the liner 20 has been completed, the air may be removed by releasing the pressure within the pressurization chamber 222

5 allowing the flexible member 220 to reconfigure into its opened domed shape and to release the seal 224 from the sealing lip 228. This may then allow for the exhaust from the liner interior to flow through the exhaust lumen 212, through an exhaust chamber 216, and further into an exhaust lumen 218 for venting from the treatment assembly, as described herein.

10 [0117] LINER REMOVAL

[0118] In further facilitating a treatment procedure, the liner may also be configured to aid in its removal from the underlying tissue after a cryoablation treatment. After the tissue has been treated, the liner 20 may remain frozen on the underlying uterine tissue preventing removal of the liner 20 from the patient's body for up to several
15 minutes. The liner 20 may be left in the patient for a period of time after the cryoablation treatment until the tissue thaws as pulling the liner 20 from the tissue prematurely may tear the liner 20; however, leaving the liner 20 in place may increase patient discomfort. Hence, to facilitate removal of the liner 20 from the underlying frozen tissue, a number of different warming techniques may be optionally implemented.

20 [0119] Circulating a warm or room temperature fluid within the liner is one method for thawing the liner 20 and adjacent tissue to expedite the removal of the liner 20. A gas (e.g., air, expanded helium, etc.) may be used instead of a liquid as a warming gas may prevent the creation of a solid which could potentially block the exhaust gas pathway. Additionally, use of a warming gas may also slow the boiling-off of any
25 remaining cryoablative liquid as boiling-off the cryoablative liquid too quickly could create a pressure spike within the liner 20. A liquid with a freezing point lower than the boiling point of the cryoablative agent, such as nitrous oxide, may be utilized but may not be required. It may also be possible to use a liquid which has a much higher boiling point than the cryoablative fluid provided that all of the cryoablative fluid has previously
30 boiled-off. Closing the actuatable valve within the exhaust block and measuring the pressure inside the liner is one way to detect if any of the cryoablative fluid remains: an increase in pressure would indicate the presence of liquid cryoablative fluid still boiling-off.

[0120] One variation is shown in the partial cross-sectional top view of Fig. 10 which shows a liner having a warming fluid lumen 230 integrated with the liner 20 and cooling probe 22. The warming fluid lumen 230 may be formed integrally with the distal portion of the liner 20 and extend proximally through the interior of the liner 20 and/or cooling probe 22. During the cryoablation treatment procedure, the warming fluid lumen 230 may remain in a flattened configuration which is non-obstructive to the introduced cryoablative agent. However, once the cryoablation treatment has been completed and the liner 20 is to be removed from the uterus UT, a fluid 232 (e.g., saline, water, etc.) which may be warmed may be introduced into the warming fluid lumen 230 from outside the patient body such that the fluid 232 flows through the lumen 230 and the liner 20 interior and out the distal portion of liner 20 through a lumen opening 234 and into direct contact against the tissue surface and the exterior of the liner 20. The fluid 232 may warm the contacted frozen tissue and facilitate the release of the liner 20 from the tissue surface as the fluid 232 pushes the liner 20 away from the frozen tissue. The fluid 232 may be introduced continuously or intermittently (e.g., via a syringe, pump, or by the treatment assembly itself) while the liner 20 is retracted from the uterus UT and out of the patient body. For fluid circulation external to the liner 20, a liquid may be used over a gas because of the relatively higher heat transfer rate which is possible with a liquid.

[0121] Figs. 11A and 11B illustrate one variation for fabricating the liner 20 with an integrated fluid lumen 230. The liner 20 may first be formed to include a lumen 230 which protrudes distally from the distal end of the liner 20, as shown in the top view of Fig. 11A. Once such a liner has been formed, e.g., via RF welding, the fluid lumen 230 may be inverted such that it passes through the interior of the liner 20, as shown in the top view of Fig. 11B. This variation illustrates one example for incorporating a fluid lumen with the liner 20; however, any number of other methods may be utilized as well.

[0122] Aside from forming an integrated fluid lumen into the liner, other mechanisms may instead be utilized to facilitate liner removal from the contacted tissue. Figs. 12A and 12B illustrate top and side views, respectively, another variation of a liner 20 incorporating a tether or wire 240 (e.g., Kevlar, Nylon, etc.) which may be positioned externally of the liner 20 such that it lies between the surrounding tissue and liner exterior. Initially, the tether or wire 240 may be loosely looped over the liner 20 to prevent any interference from the tether or wire 240 for liner deployment. Alternatively, the tether or wire 240 may be integrally formed with the liner 20 material itself or the tether or wire 240 may be attached along the interior surface of the liner 20. In either

case, the tether or wire **240** may be attached at shaft attachment points **242** while extending over the entire length of the liner **20** when deployed against the tissue.

[0123] Once the liner is ready to be removed from the uterine tissue, the tether or wire **240** may be tensioned either via a control mechanism on the device handle or via simply pulling on the device. The variation shown in the top and side views of Figs. 13A and 13B illustrate how retraction of the sheath **12** may tension the tether or wire **240** such that the liner **20** may begin to pull away **244** from the tissue. The presence of the tether or wire **240** may help to ensure that the liner **20** remains attached to the treatment assembly. The distance **246** that the liner **20** has stretched during sheath retraction **12** may be seen in Fig. 13A. The tether or wire **240** may continue to be tensioned until the liner **20** has pulled entirely from the tissue surface.

[0124] In yet another variation, Fig. 14 shows a top view of a liner **20** which may incorporate a guide hole **250** at a distal end region of the liner **20**. The tether or wire **240** may pass externally of the liner **20** into and through the guide hole **250** such that the tether or wire **240** may remain looped around the liner exterior.

[0125] Fig. 15 shows yet another variation where a guide tube **252** may be integrally formed with the liner material either externally or internally of the liner **20**. The guide tube **252** may extend laterally along the distal portion of the liner **20** such that the tether or wire **240** may be looped securely through the guide tube **252**.

[0126] Fig. 16 shows yet another variation where two or more guide holes **254** may be formed along the interior of the liner **20**. The tether or wire **240** may be attached to a secondary tether or directly to each of the guide holes **254**.

[0127] Fig. 17 shows yet another variation where a peripheral channel **256** may be formed around the periphery of the liner **20** such that the tether or wire **240** extends through the peripheral channel **256** entirely around the liner **20**.

[0128] Fig. 18 shows yet another variation where the two or more guide holes **254** formed along the liner interior with a tether **258** extending between the holes **254** may also be attached to the probe **22**. Retraction of the probe **22** may tension the distal end of the liner **20** to facilitate its release from the tissue.

[0129] In these examples and any of the variations herein, a wire or heating element which may be warmed or energized (e.g., infrared) may be located on the probe shaft or positioned within the liner **20** interior. Once the treatment procedure has been completed, the wire or heating element may be activated to warm the liner **20** and the adjacent contacted tissue to facilitate the thawing of the tissue for removal of the liner **20**.

[0130] Additionally and/or alternatively, the liner 20 may be comprised of a lubricious liner or a separate non-stick coating may be applied to the liner exterior. It is not uncommon for polymers such as urethanes, especially thin films, to stick together if tightly-packed during sterilization, transportation and storage. The liner 20, being a thin polyurethane film compressed into a sheath, may employ a lubricious material or surface to ensure that the liner 20 fully deploys and inflates following unsheathing. For instance, the lubricity of the liner 20 may be increased through the formulation of the urethane blend used to make the thin film. Diatomaceous earth may also be utilized as an additive to the liner material to make the surface irregular to prevent the film from sticking to itself. Lubricants can also be added to the interior and/or exterior of the liner 20 to increase its lubricity and prevent the liner from sticking to itself after being sheathed. Silicone oil and talc are examples of two possible lubricants. Surface lubricants can also be applied to the film before or after the liner is RF-welded.

[0131] Another method for preventing the liner from sticking to itself is to keep the liner 20 unsheathed until just prior to delivery and deployment. A simple liner folding tool can be included within the sterile package to enable the user to easily load the liner 20 into the sheath in its pleated configuration. In order to prevent the device from being used prior to the sheath being advanced over the liner 20, the position of the sheath may be used as an input in a system check algorithm.

[0132] Because optimal ablation coverage and depths may not be uniform over the entire contacted tissue region, the liner 20 may be adjusted in thickness over particular regions of the liner 20 to insulate predetermined tissue regions to result in tailored ablations. Ablation depths may be shallower where the liner 20 is relatively thicker due to less efficient thermal transfer across the thicker areas. One example is shown in the top view of Fig. 19 which illustrates the liner 20 having insulated regions 260 of the liner (e.g., thicknesses greater than 0.0012 in.), for instance, near the distal portions of the liner such as near the uterine cornua and lower segment when the liner 20 is deployed within a uterine cavity. Alternatively, multiple layers of the membrane may be utilized where liner thickness is to be increased. Having the thickened regions 260 insulate the contacted tissue may help to prevent intrauterine adhesions. Another variation may include a liner 20 having, for instance, quilted pockets filled with air or gas to insulate targeted regions and reduce ablation depth and possibly even coverage.

[0133] In yet another variation, the liner 20 may be designed with one or more predetermined weak points, as shown in the top view of Fig. 20. If excessive tension

were applied to the liner **20** while it is frozen to tissue, the liner **20** could tear. By locating one or more weakened regions **270** of the liner **20** near, e.g., the proximal connection to the probe shaft, the liner **20** may be designed to tear specifically at the designated weakened regions **270** which may make retrieval of the detached liner **20** as a single piece relatively easier once the tissue fully thaws. It is preferable that the liner **20** remain intact when tension is applied during the removal of the device from the patient body. However, it may be advantageous to have designated weakened regions **270** to prevent too much force from being applied to the uterine tissue. The location of the weakened regions **270** may also be chosen to facilitate removal of the liner **20** as a single piece after the uterus has warmed following a cryotherapy treatment.

[0134] In yet another variation of the liner, the liner may be separated into several individual liners in a multi-liner assembly, as shown in the variation of Fig. 21. Because it is not uncommon for uterine cavities to be arcuate, septate, bicornuate or have other abnormalities, the liner may be configured to have two more liners which could be deployed simultaneously. Such a multi-liner probe may deliver different cryotherapy or hyperthermic therapies to different parts of the uterus, if so desired. In the variation shown, a single probe shaft **280** may include a first cooling probe **282** and a second cooling probe **284** extending at an angle from the probe shaft **280**. The cooling probes **282**, **284** may each have a respective first liner **286** and second liner **288** where each liner may be deployed to function in the same manner as described herein. Moreover, while two separate liners are shown, other variations may include more than two individual liners depending upon the desired treatment results.

[0135] PROBE CONTROL

[0136] In any of the variations described herein, the cooling probe **22** may optionally include a compressible tip **290** having a collapsible opening **292** defined through the tip **290**, as shown in the top view of Fig. 22A. The compressible tip **290** may be positioned upon the distal tip of the probe **22** located within the interior of the liner **20**. Because the probe **22** may be translatable within the liner **20** and relative to the sheath **12**, the tip **290** may present a soft and atraumatic surface in the event the probe **22** is advanced into contact against the interior of the liner **20** and underlying tissue surface, as shown in Fig. 22B, to prevent liner tears or trauma to the uterine tissue **UT**. If the tip **290** is compressed against the liner **20** and/or tissue, the opening **292** may expand laterally to increase in diameter. In alternative variations, any number of relatively soft materials

which can withstand the temperature of the cryoablative agent may be utilized and other shapes and structures may also be utilized.

[0137] Aside from the tip contacting the interior of the liner **20**, the cooling probe **22** itself may become inadvertently buried or urged into the anterior or posterior tissue surface of the uterus **UT** when deployed within the uterine cavity. This is due to the orientation of the uterus **UT** which is typically angled relative to the vaginal opening of the patient (anteverted or retroverted) as well as along the body of the uterus **UT** itself (anteflexed or retroflexed), as shown in the cross-sectional side view of Fig. 23. With the sheath **12** introduced through the cervix **CV** and retracted and the liner **20** deployed and expanded into contact within the uterine cavity, the cooling probe **22** may have a tendency to be urged into the posterior tissue wall of the uterine cavity when advanced or adjusted into position within the liner interior, as illustrated. However, the cooling probe **22** is preferably located at a central position within the expanded liner interior, as illustrated, in order to allow for the uninhibited infusion of the cryoablative fluid from the infusion line **52**. Having the probe **22** placed directly against the liner interior and underlying tissue wall may obstruct the infusion line **52** potentially resulting in an uneven ablation pattern in the tissue.

[0138] Hence, the cooling probe **22** may be fabricated from a material such as annealed stainless steel which may provide the probe **22** with improved ductility due to its minimized internal stresses. This improved ductility allows for the probe **22** to flex relative to the sheath **12** away from the walls of the liner interior and contacted tissue walls such that the probe **22** may be centered within the expanded liner **20** particularly in the reduced temperature environment during cryoablation. The use of fully annealed stainless steel may facilitate the positioning of the probe **22** within the liner **20** due in part to the slots defined along the probe **22** but also due to the material properties. For instance, the probe **22** fabricated from annealed stainless steel may have, e.g., a 0.006 in. wall thickness, selected to minimize overall device diameter while maintaining sufficient hoop-strength to prevent it from being crushed or kinked in-use. Annealing the slotted stainless steel exhaust tube may also improve its cyclic fatigue life compared to a half-hard or a full-hard material.

[0139] In yet other variations, rather than utilizing annealed stainless steel, other shape set materials may be used. For instance, a simple steel cannula or a more complex articulating cannula which may incorporate multiple individual elements which are flexible in a first state and optionally locked into a particular configuration in a second

state may also be used although not necessary. However, it is desirable that the cannula takes the shape of the uterine cavity within the sagittal plane of the patient and uterus **UT** without being biased to either the anterior or posterior side of the uterus **UT**. Provided that the cannula flexes with the shape of the uterus **UT**, holds its flexed position during the cryoablation treatment, and also has substantial radial strength to prevent crushing and/or kinking of its shape, any number of shape set materials may be utilized.

[0140] RESERVOIR TEMPERATURE CONTROL

[0141] During a cryotherapy treatment, it is desirable to control the amount of the cryoablative agent delivered into and through the liner **20**. A few of the parameters which may affect the flow rate and volume of the cryoablative agent discharged from the reservoir or canister **92** may include temperature of the treatment assembly **80** and reservoir or canister **92** as well as ambient temperature in which the assembly **80** is used as such temperatures can affect the internal pressure of the reservoir or canister **92**. Unless controlled by other methods, the flow rate of a pressurized cryogen is generally controlled by the internal pressure of the vessel in which the cryoablative agent is contained. Thus, in order to be able to deliver a consistent cryoablative therapy over a range of device and room temperatures, controlling either the inflow rate or starting temperature and pressure of the cryogen is desirable.

[0142] One method for controlling the starting cryogen pressure is by designing the system to operate at the high end of the temperature range and heating the reservoir or canister **92** to a specified temperature and corresponding internal pressure. The heat could be supplied by a various mechanisms such as an electrical heating element wrapped around the reservoir or canister **92**. In one variation, the electrical power for the heating element could be provided by a battery within the device itself.

[0143] In another variation, the electrical power may be provided by a heating cradle **316** prior to device use, as shown in the exemplary schematic side view of Fig. 24. A separate warming cradle **316** may define a receiving channel **318** sized to receive the reservoir housing **88** of the treatment assembly **80**. The cradle **316** may further include an electrical connector **320** connected to an optional power supply (DC) **322** (which may be recharged) and/or the cradle **316** may be electrically connected to a stationary power supply via a power supply (AC) line **326**. The cradle **316** may also incorporate an optional stabilizing weight **324** to provide for stability when the treatment assembly **80** is docked within the receiving channel **318**.

[0144] The treatment assembly **80** itself may incorporate a heating element **300** (e.g., a resistive heating element) which may be wrapped partially or entirely around the reservoir or canister **92**. A layer of insulation **304** may also be provided around the reservoir or canister **92** to provide for a thermally stabilized warming environment. A temperature sensor **302** (e.g., thermocouple, thermistor, etc.) may also be incorporated for thermal contact with the reservoir or canister **92** for sensing the canister temperature. An electrical connector **314** may be located correspondingly along the reservoir housing **88** for electrically contacting the electrical connector **320** positioned upon the cradle **316** such that the cradle **316** may provide electrical power to the treatment assembly **80** when docked within the cradle receiving channel **318**.

[0145] A connection sensor **312** may be electrically connected to the connector **314** such that the sensor **312** may provide a signal via connection line **310** to the microprocessor or controller **110** indicating that the treatment assembly **80** has been docked and is able to receive power from the cradle **316**. The heating element **300** may also be electrically coupled to the microprocessor or controller **110** via heating element line **306** and the temperature sensor **302** may likewise be coupled to the microprocessor or controller **110** via temperature sensor line **308**.

[0146] With the heating element **300** and temperature sensor **302** so coupled to the microprocessor or controller **110**, the heating assembly may form a closed-loop system where the microprocessor or controller **110** may be programmed via a software algorithm to control the electrical power supplied to the heating element **300** depending upon the measured temperature of the temperature sensor **302** such that the reservoir or canister **92** is heated to a predetermined temperature or maintained within a predetermined temperature range prior to a cryotherapy treatment. The insulation **304** may accordingly slow the rate of cooling of the reservoir or canister **92** and also extend the available time between the removal of the treatment assembly **80** from the cradle **316** and the initiation of the treatment.

[0147] The microprocessor or controller **110** may be further programmed to alert or indicate (e.g., auditory or visual) to the user that the treatment should be initiated before the sensed temperature (and pressure) of the reservoir or canister **92** drops below a set point temperature desired for completion of a full cryotherapy treatment. An auditory and/or visual indicator (e.g., lights, alarms, or other visual or auditory cues) may also be incorporated to the device assembly **80** and/or cradle **316** to indicate to the user that the device assembly **80** is actively heating or has reached its target temperature or pressure.

In yet another variation, the microprocessor or controller **110** controlling the heating of the reservoir or canister **92** can be located in the cradle **316**. The reservoir or canister **92** or pressure input would have to be supplied via a direct electrical connection or a wireless connection between the cradle **316** and device assembly **80**.

5 **[0148]** In yet another variation for maintaining the reservoir or canister **92** at a predetermined temperature, one or more heating elements may instead be located in the cradle **316** and used to transfer thermal energy to the reservoir or canister **92**. The microprocessor or controller **110** (or an additional microprocessor) may be located in the cradle **316** in electrical communication with the one or more heating elements.

10 **[0149]** In yet another variation, the cradle **316** may be configured to both heat and cool the reservoir or canister **92**. A thermoelectric unit (e.g., Peltier device) or other source of refrigeration (e.g., cooler, ice bath, etc.) may be used to cool the reservoir or canister **92** to the desired temperature range. In the case of a thermoelectric unit, such a device may be used to both cool or heat the reservoir or canister **92**. In either case, the
15 various components of the closed-loop control system could be located either in the device **80** or cradle **316** as described herein.

[0150] While specific variations are described, it is intended that each of the features described above may be combined in any number of different combinations and such combined features are intended to be within the scope of this disclosure. For
20 instance, the treatment assembly **80** shown in Figs. 4A-4C may incorporate each of the features such as the reservoir or canister valve **94** and modulation control unit **96** (as shown in Figs. 5 and 8) as well as the liquid exhaust trap **130** (as shown in Fig. 6) and exhaust collectors **150** or **160** (as shown in Figs. 7A and 7B) into a single embodiment. The treatment assembly **80** may also incorporate the valve located within the exhaust
25 block **46** (as shown in Fig. 9) as well as any of the liner removal variations as shown in Figs. 10 to 18. Furthermore, the features of Figs. 19 to 22B may also be incorporated into a single embodiment as well. The additional feature of a probe **22** or device utilizing annealed stainless steel (as shown in Fig. 23) may also be incorporated into the single embodiment as well. Additionally, any embodiment incorporating any of the various
30 combinations may be optionally utilized with the features of a reservoir temperature control assembly and cradle **316** (as shown in Fig. 24).

[0151] While illustrative examples are described above, it will be apparent to one skilled in the art that various changes and modifications may be made therein. Moreover, various apparatus or procedures described above are also intended to be utilized in

combination with one another, as practicable. The appended claims are intended to cover all such changes and modifications that fall within the true spirit and scope of the invention.

CLAIMS

What is claimed is:

1. An inflow reservoir or canister valve, comprising:
a valve body;
5 a reservoir interface extending from the valve body and configured for fluidly coupling with a reservoir or canister containing a cryoablative agent;
a modulation control interface defined along the body and configured for fluidly coupling to a modulation control coupler;
a valve stem seated within a valve stem channel defined within the valve body;
10 an inflow lumen defined through the valve body and extending between the reservoir interface and the modulation control interface, where the valve stem is movable between a first position which obstructs the inflow lumen and a second position which opens the inflow lumen; and
a venting lumen defined through the valve body and extending between the
15 reservoir interface and a vent opening.
2. The valve of claim 1 further comprising a reservoir lumen insert extending from the reservoir interface.
- 20 3. The valve of claim 1 further comprising a vent piston which is movable between a first position which obstructs the venting lumen and a second position which opens the venting lumen.
4. The valve of claim 3 further comprising a removable locking mechanism
25 which is positionable to obstruct movement of the vent piston.
5. The valve of claim 1 further comprising a reservoir engagement control which is coupled to the valve stem, wherein actuation of the reservoir engagement control moves the valve stem from its first position to its second position.

30

6. The valve of claim 5 further comprising a microprocessor in electrical communication with the reservoir engagement control such that actuation of the reservoir engagement control is indicative to the microprocessor of an open reservoir or canister.

5 7. The valve of claim 1 further comprising a modulation control unit engaged with the modulation control interface.

8. The valve of claim 7 wherein the modulation control unit comprises a actuatable solenoid valve.

10

9. The valve of claim 1 further comprising a microprocessor in electrical communication with the inflow reservoir or canister valve.

10. The valve of claim 9 further comprising one or more pressure or temperature
15 sensors in communication with the microprocessor.

11. The valve of claim 1 further comprising:
an elongate probe having a distal tip and a flexible length;
at least one infusion lumen positioned through or along the elongate probe and in
20 fluid communication with the inflow lumen, wherein the infusion lumen defines one or
more openings along its length; and
a liner expandably enclosing the probe.

12. The valve of claim 11 further comprising at least one delivery lumen slidably
25 positioned through or along the infusion lumen, wherein translation of the delivery lumen
relative to the infusion lumen controls a number of unobstructed openings along the
infusion lumen.

13. The valve of claim 11 further comprising an exhaust lumen in fluid
30 communication with the elongate probe.

14. The valve of claim 13 further comprising a liquid exhaust trap in fluid
communication with the exhaust lumen.

15. The valve of claim 13 further comprising an exhaust collector in fluid communication with the exhaust lumen.

5 16. The valve of claim 13 further comprising an actuatable valve in fluid communication with the exhaust lumen.

10 17. The valve of claim 11 further comprising a fluid lumen defined through an interior of the liner while fluidly isolated from the interior and where the fluid lumen is in communication with an exterior of the liner.

18. The valve of claim 11 further comprising at least one wire or tether attached to the liner, wherein tensioning of the wire or tether retracts the liner.

15 19. The valve of claim 11 further comprising an atraumatic tip attached to a distal end of the probe.

20. The valve of claim 11 further comprising a cradle which is configured to receive a handle assembly enclosing the reservoir or canister.

20 21. The valve of claim 1 further comprising a warming element in thermal communication with the reservoir or canister.

22. A tissue treatment system, comprising:
an elongate probe having a distal tip and a flexible length;
25 at least one infusion lumen positioned through or along the elongate probe, wherein the infusion lumen defines one or more openings along its length;
a liner expandably enclosing the probe;
an inflow reservoir or canister valve fluidly coupled with a reservoir or canister containing a cryoablative agent;
30 a modulation control unit fluid coupled with the inflow reservoir or canister valve and in fluid communication with the at least one infusion lumen; and
a warming element thermally coupled with the reservoir or canister.

23. The system of claim 22 further comprising a microprocessor in electrical communication with the inflow reservoir or canister valve and modulation control unit.

5 24. The system of claim 23 wherein the microprocessor is in electrical communication with the warming element.

25. The system of claim 22 further comprising at least one delivery lumen slidingly positioned through or along the infusion lumen, wherein translation of the delivery lumen relative to the infusion lumen controls a number of unobstructed openings
10 along the infusion lumen.

26. The system of claim 22 further comprising an exhaust lumen in fluid communication with the elongate probe.

15 27. The system of claim 26 further comprising a liquid exhaust trap in fluid communication with the exhaust lumen.

28. The system of claim 26 further comprising an exhaust collector in fluid communication with the exhaust lumen.
20

29. The system of claim 26 further comprising an actuatable valve in fluid communication with the exhaust lumen.

30. The system of claim 22 further comprising a fluid lumen defined through an interior of the liner while fluidly isolated from the interior and where the fluid lumen is in communication with an exterior of the liner.
25

31. The system of claim 22 further comprising at least one wire or tether attached to the liner, wherein tensioning of the wire or tether retracts the liner.
30

32. The system of claim 22 further comprising an atraumatic tip attached to a distal end of the probe.

33. The system of claim 22 further comprising a cradle which is configured to receive a handle assembly enclosing the reservoir or canister.

34. A method of treating tissue, comprising:

- 5 monitoring a temperature or pressure of a reservoir or canister containing a cryoablative agent;
- maintaining the temperature or pressure of the reservoir or canister at a predetermined level;
- positioning an elongate probe into a body lumen to be treated;
- 10 expanding a liner enclosing the probe into contact against the body lumen; and
- infusing a cryoablative agent through a delivery lumen such that the cryoablative agent passes into an infusion lumen, through one or more unobstructed openings, and into contact against an interior of the liner.

- 15 35. The method of claim 34 further comprising adjusting a position of the delivery lumen relative to the infusion lumen which is positioned through or along the elongate probe such that one or more openings defined along a length of the infusion lumen remain unobstructed by the delivery lumen.

- 20 36. The method of claim 34 further comprising maintaining the temperature of the reservoir or canister via a warming element in thermal communication with the reservoir or canister.

37. The method of claim 34 wherein monitoring a temperature comprises
- 25 monitoring the temperature via a microprocessor in electrical communication with a temperature sensor in thermal communication with the reservoir or canister.

38. The method of claim 34 further comprising monitoring a flow of the cryoablative agent from the reservoir or canister via an inflow reservoir or canister valve
- 30 fluidly coupled with the reservoir or canister.

 39. The method of claim 38 further comprising modulating the flow of the cryoablative agent through the inflow reservoir or canister valve via a modulation control

unit fluid coupled with the inflow reservoir or canister valve and in fluid communication with the infusion lumen.

40. The method of claim 34 further comprising passing exhaust gas from the interior of the liner into an exhaust lumen in fluid communication with the elongate probe.

41. The method of claim 40 further comprising passing the exhaust gas through a liquid exhaust trap in fluid communication with the exhaust lumen.

42. The method of claim 40 further comprising collecting the exhaust gas in an exhaust collector in fluid communication with the exhaust lumen.

43. The method of claim 34 further comprising introducing a warming liquid or gas between an exterior of the liner and a contacting tissue surface.

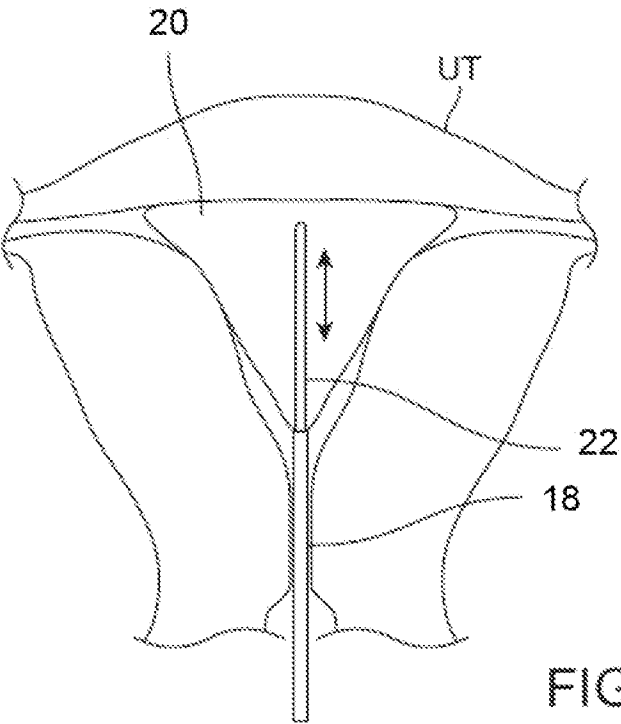
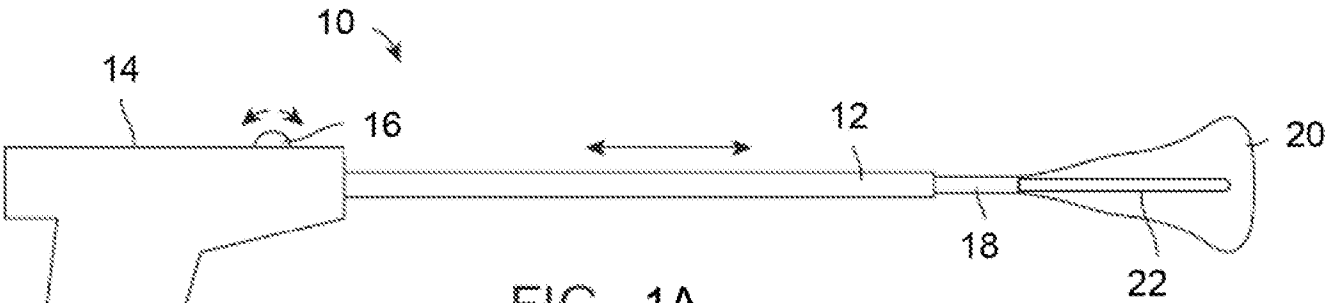
44. The method of claim 34 further comprising urging an exterior of the liner from a contacting tissue surface via a wire or tether.

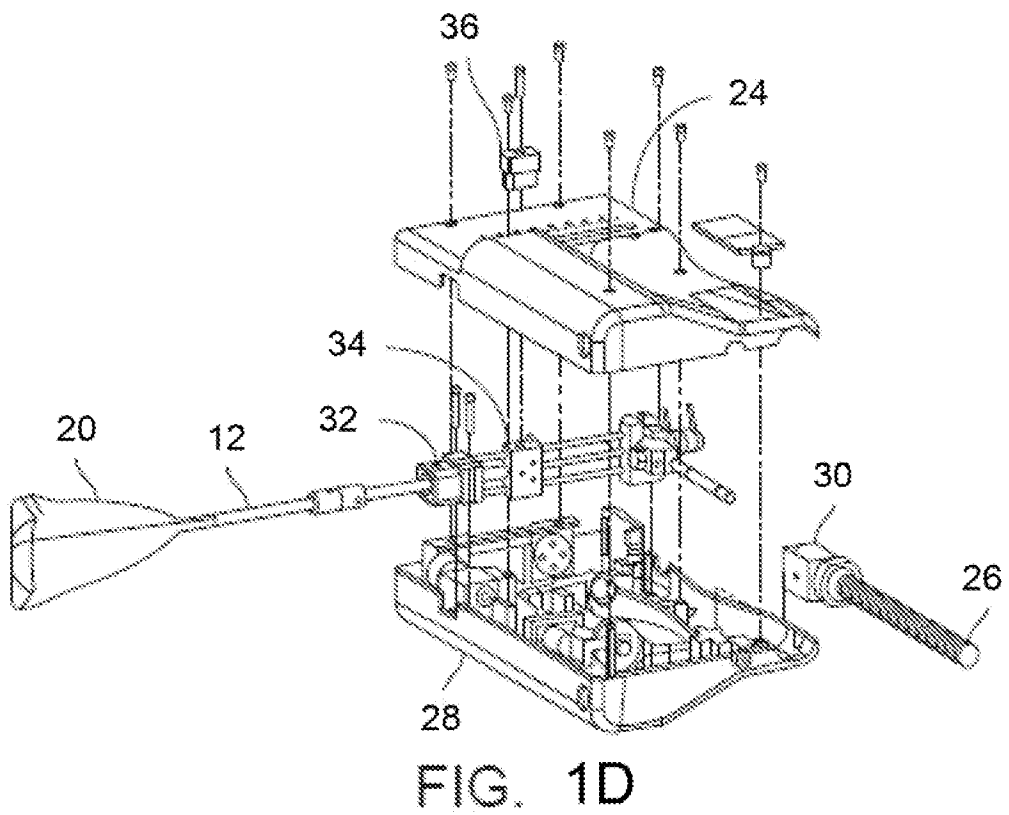
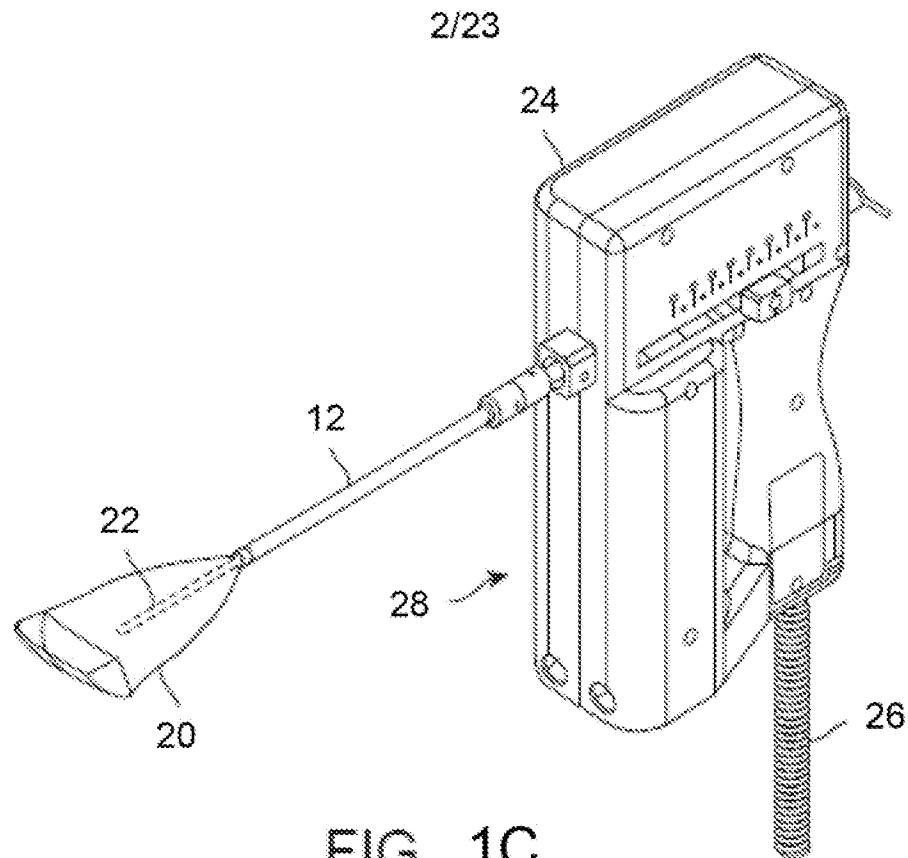
45. The method of claim 34 further comprising supporting a handle assembly containing the reservoir or canister via a cradle assembly.

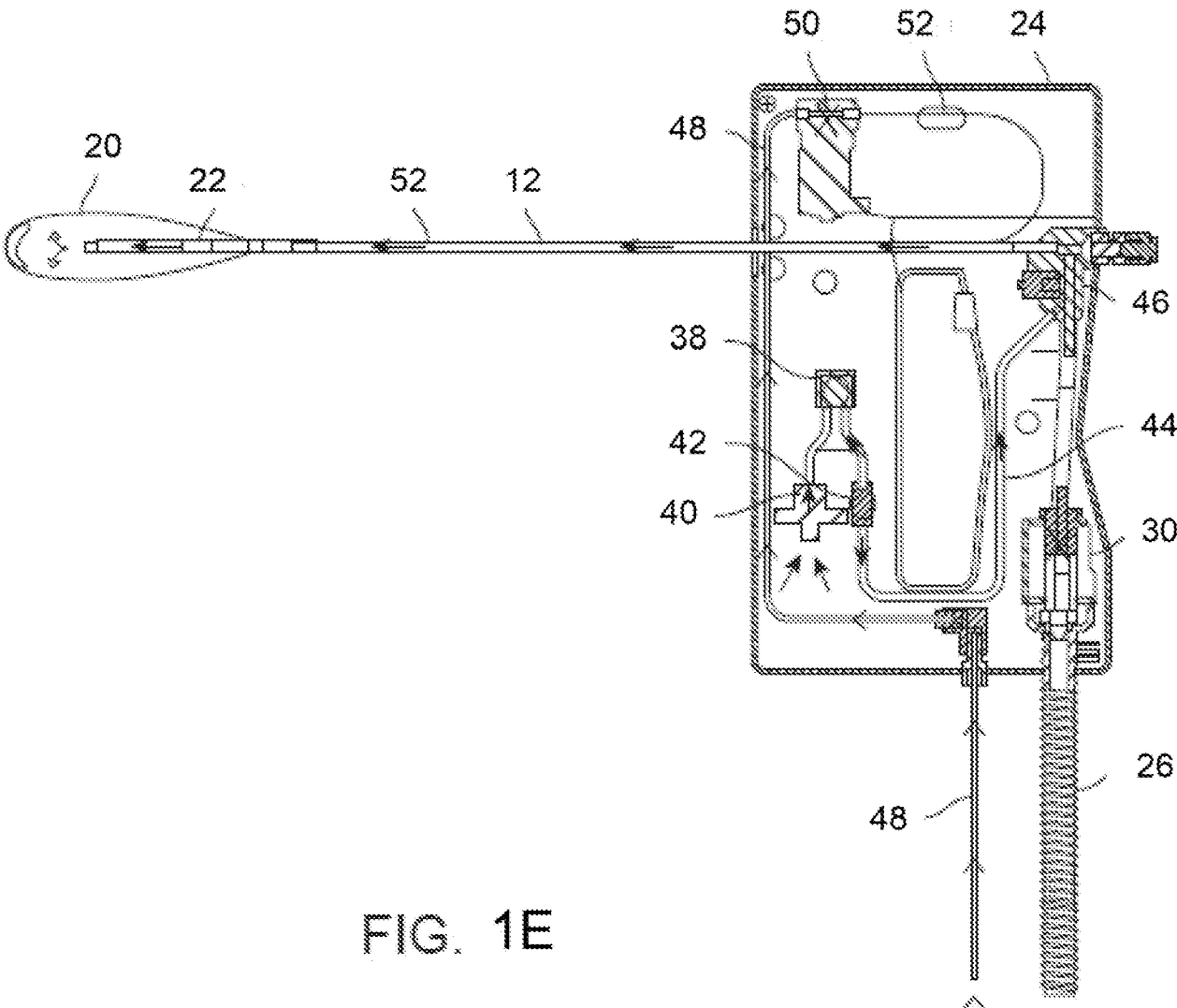
46. The method of claim 34 wherein expanding a liner comprises infusing one or more puffs of air into the liner until the liner conforms to the body lumen.

47. The method of claim 35 wherein adjusting a position of a delivery lumen comprises translating the delivery lumen through the infusion lumen to selectively obstruct the one or more openings.

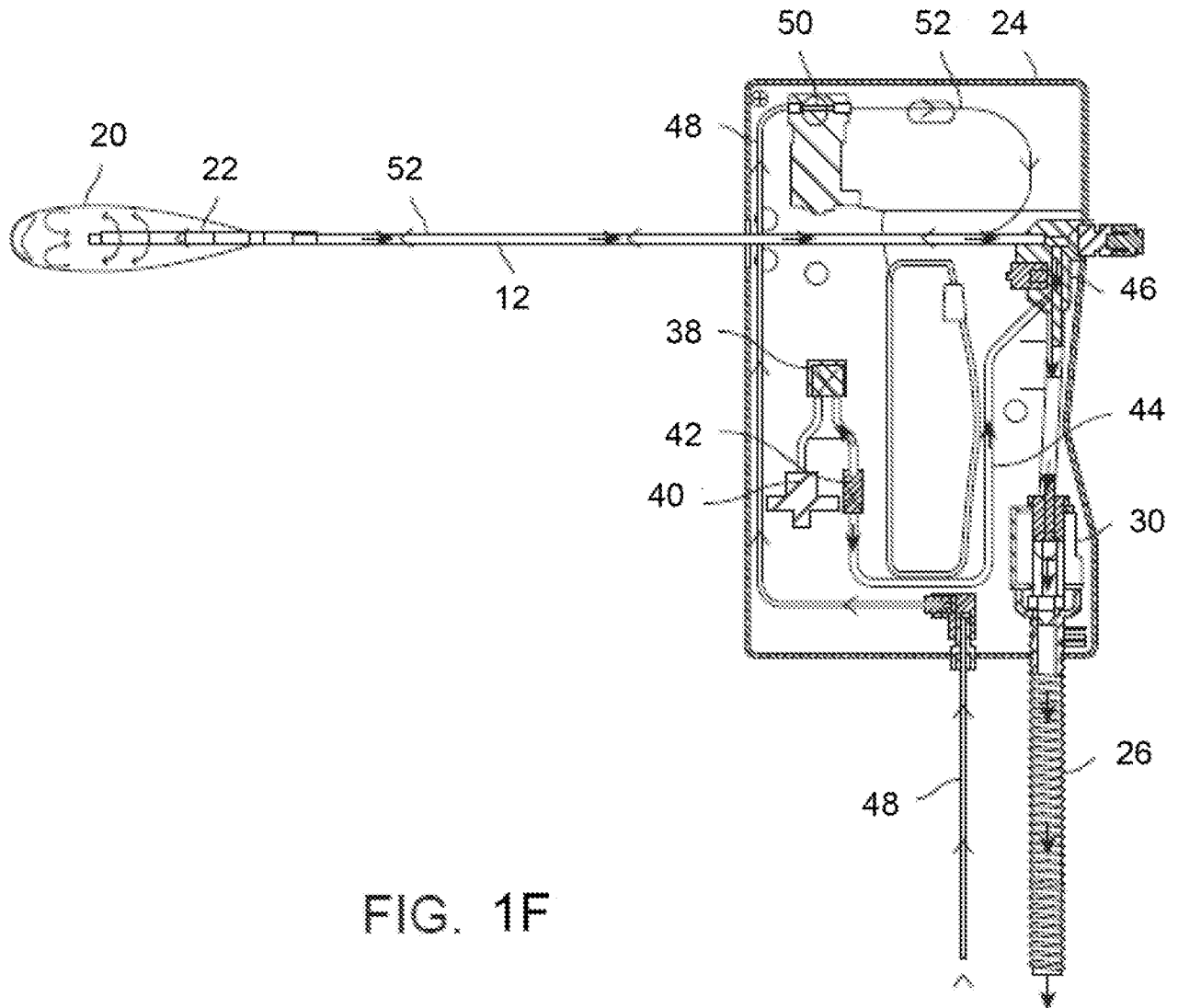
48. The method of claim 34 wherein infusing a cryoablative agent comprises infusing nitrous oxide into the interior of the liner.

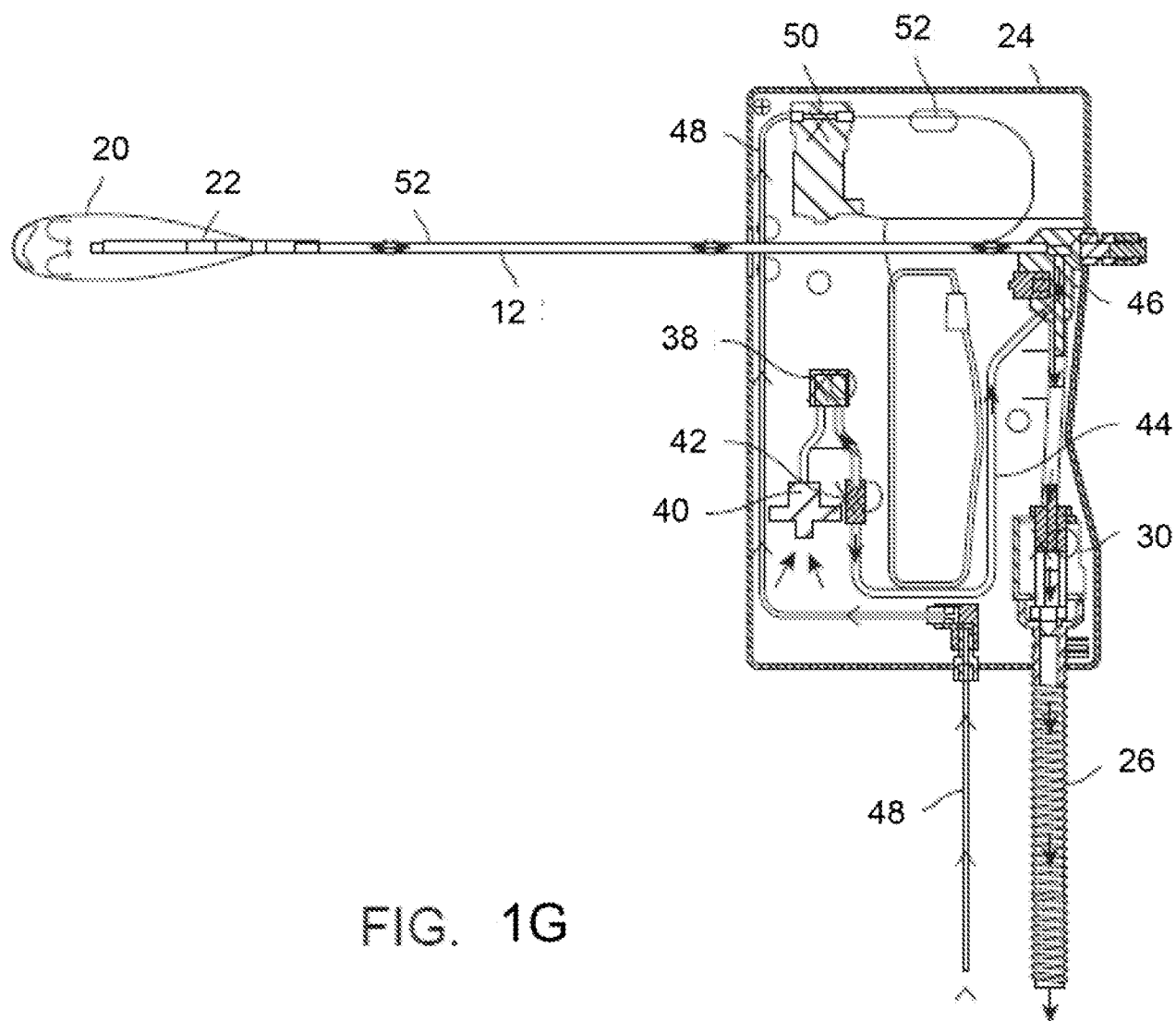






4/23





6/23

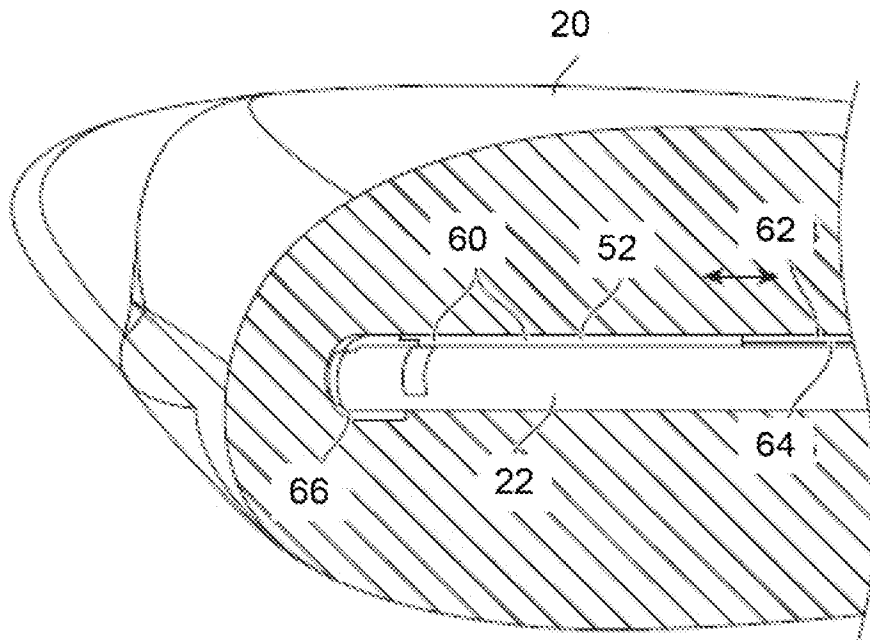


FIG. 2A

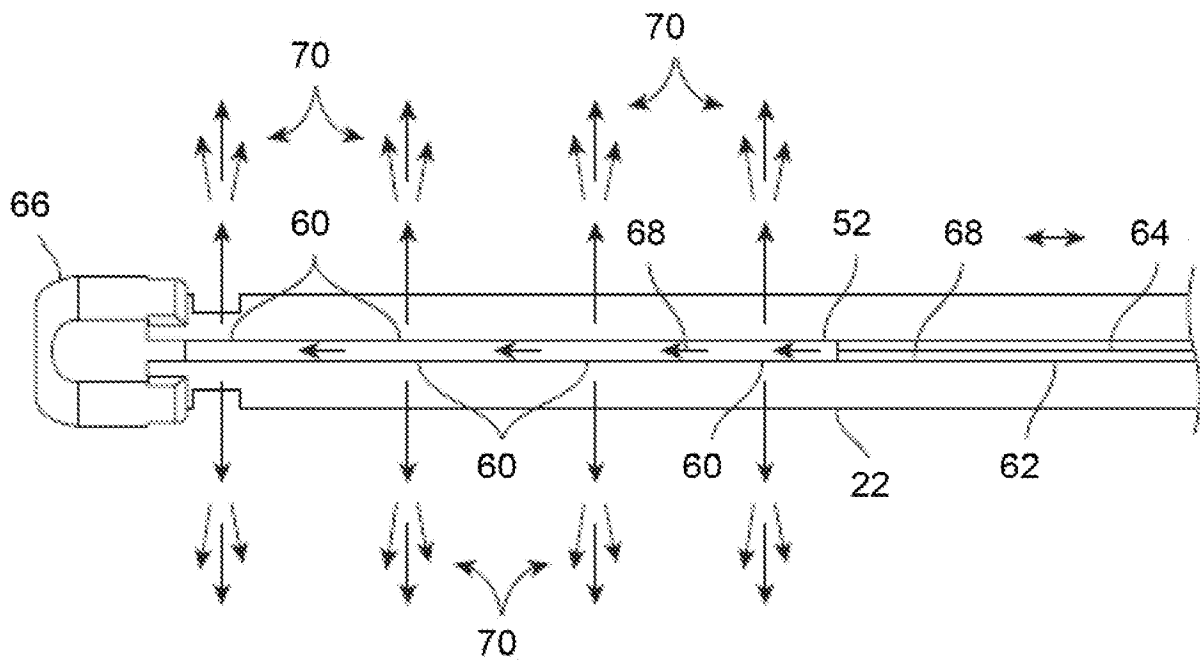


FIG. 2B

7/23

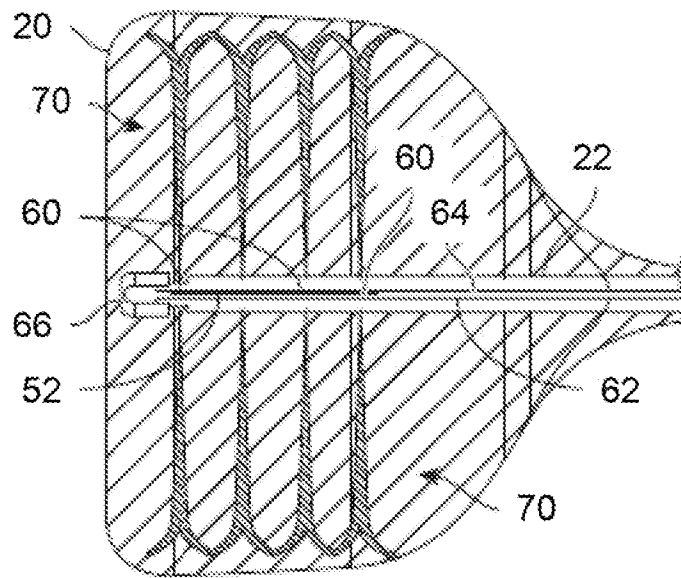


FIG. 3A

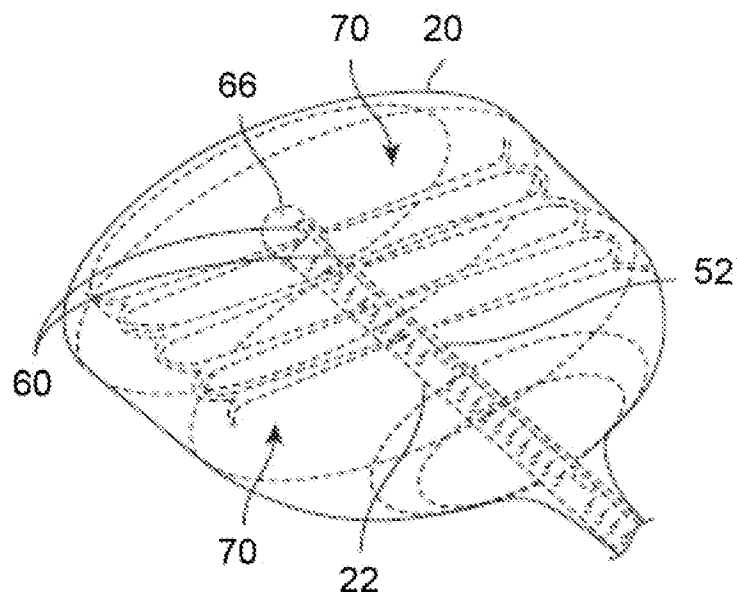
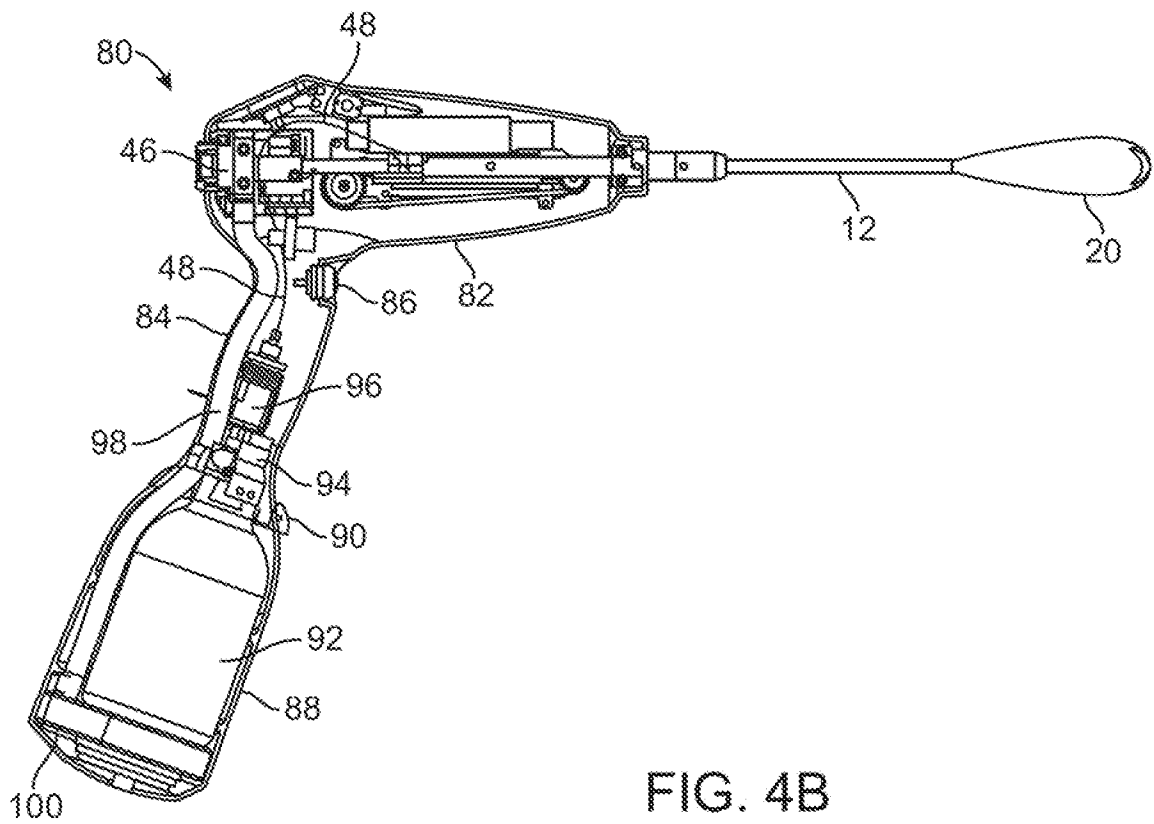
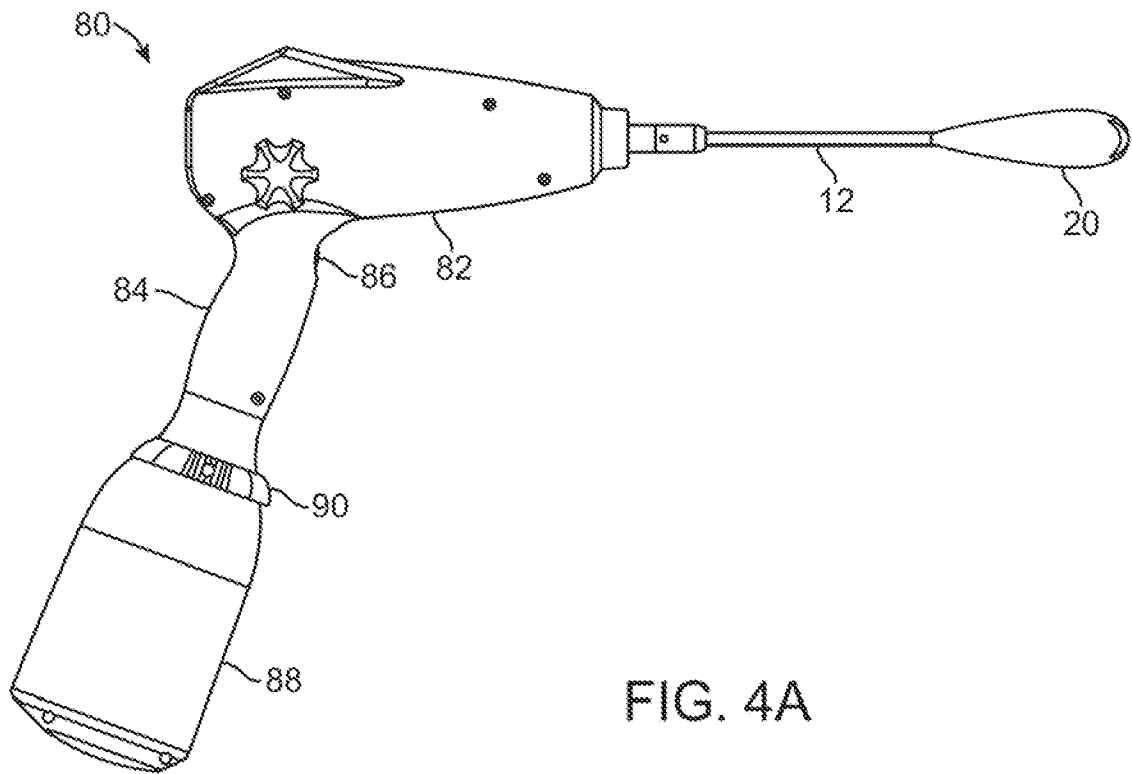


FIG. 3B

8/23



9/23

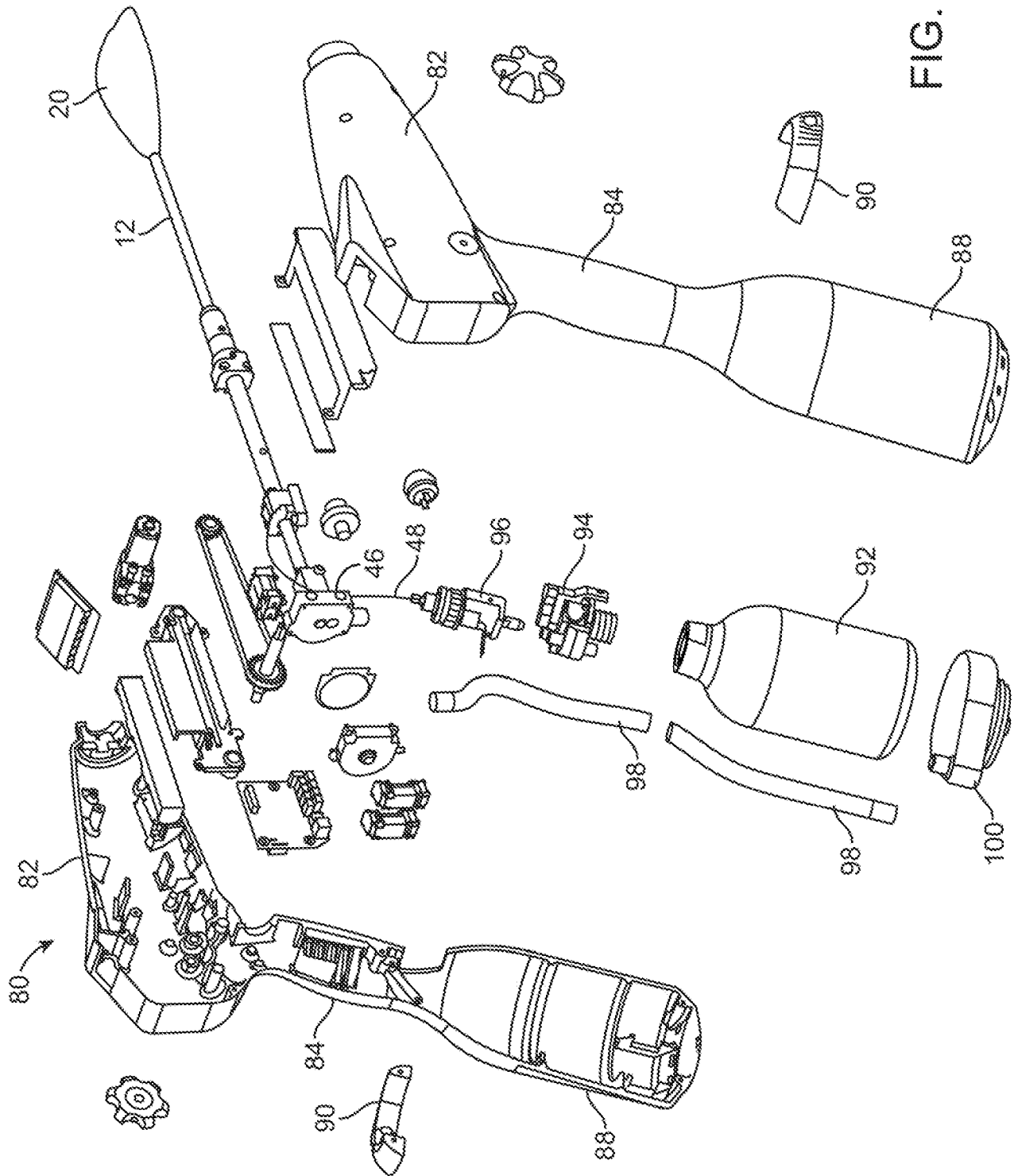


FIG. 4C

10/23

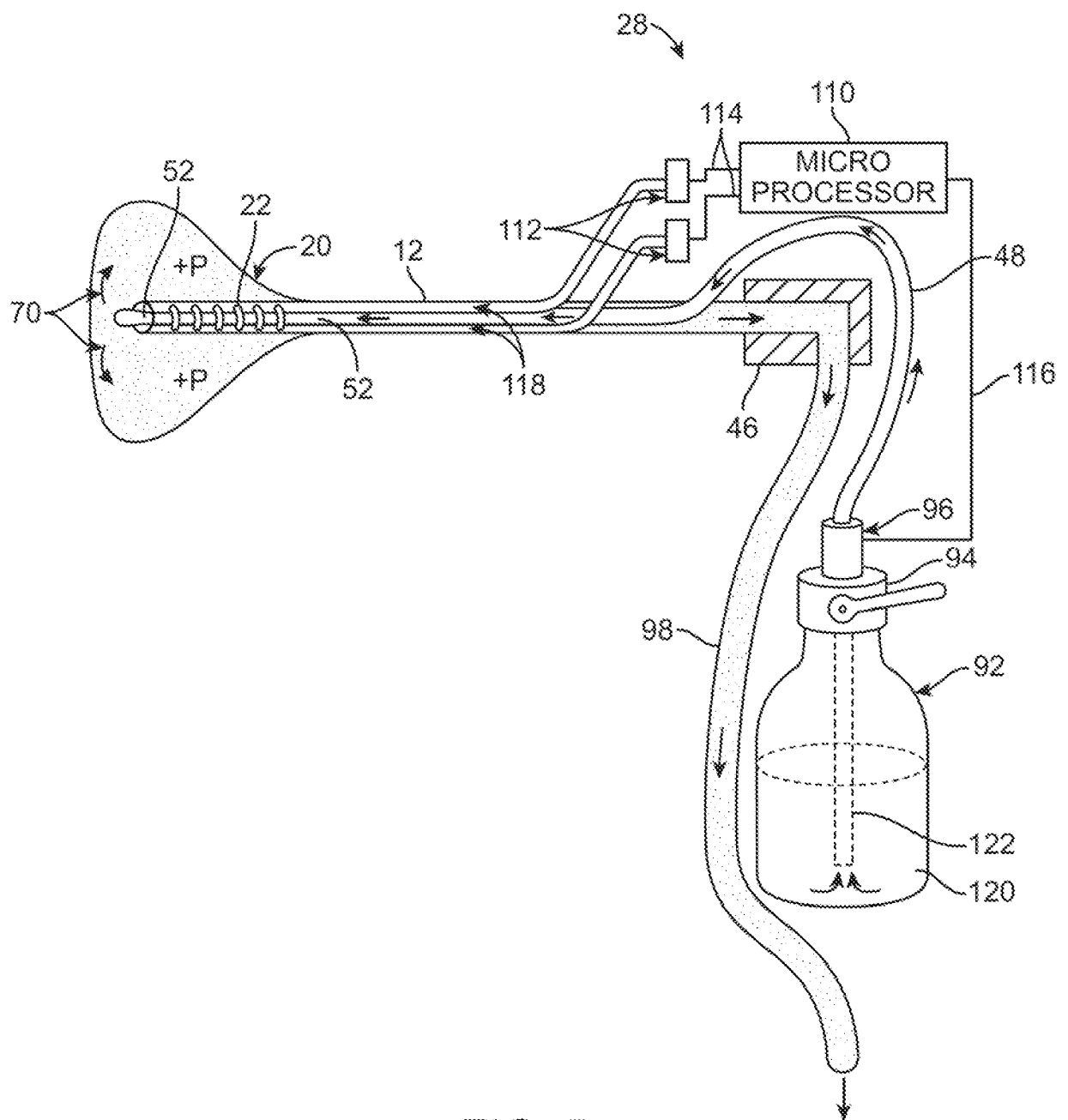
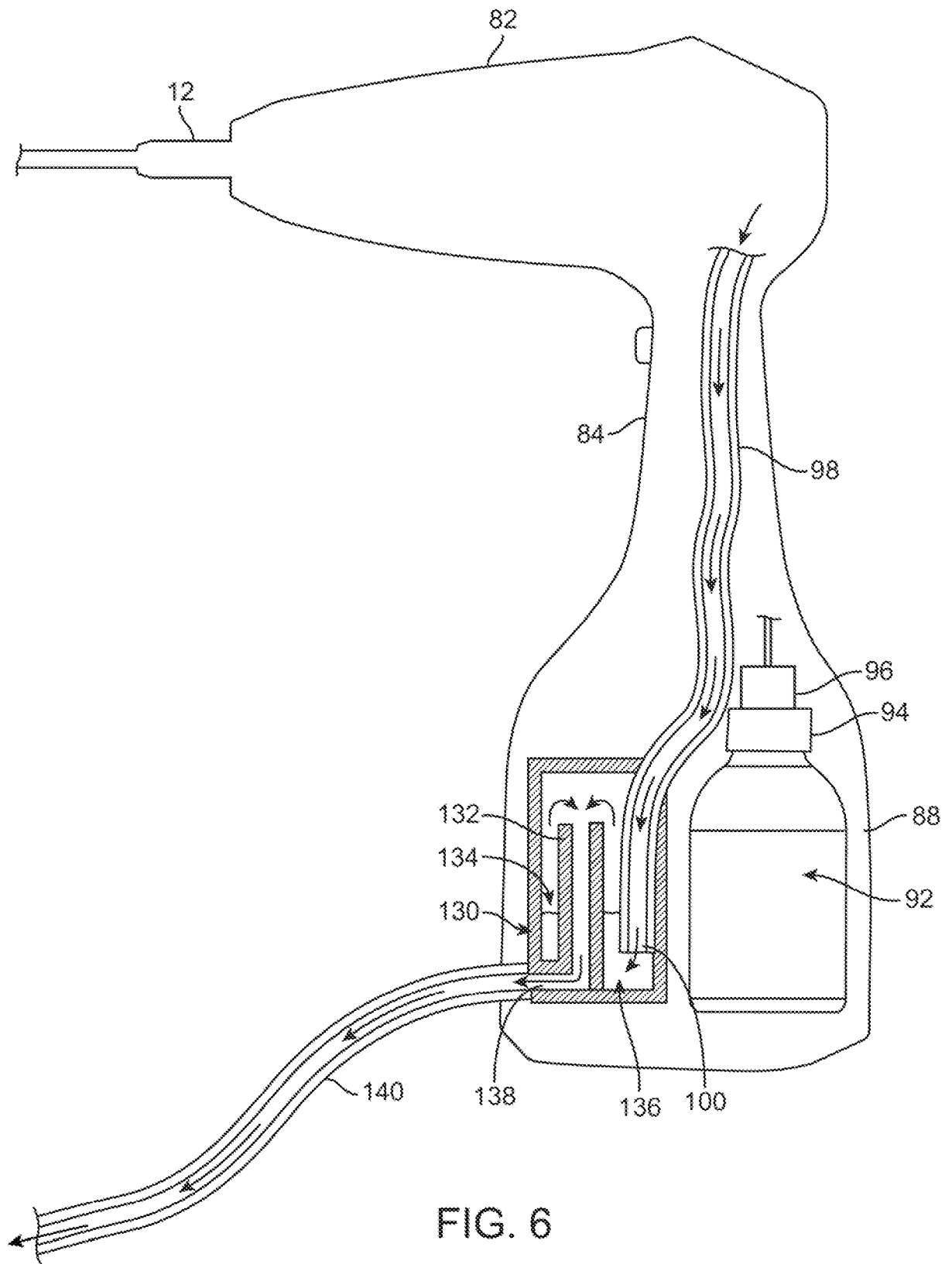


FIG. 5

11/23



12/23

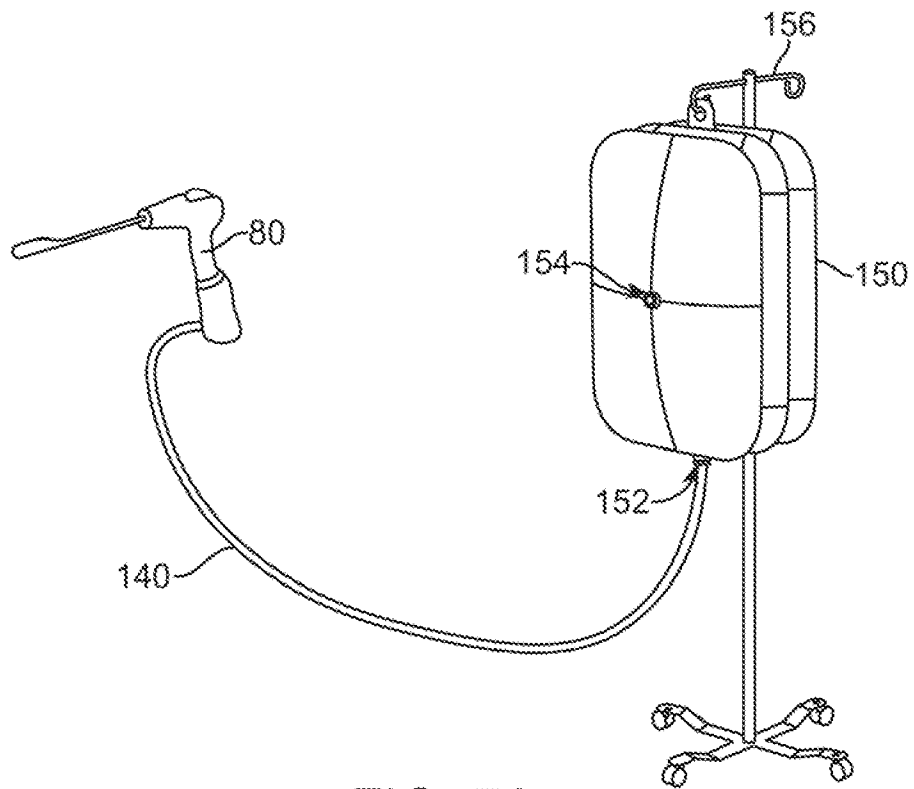


FIG. 7A

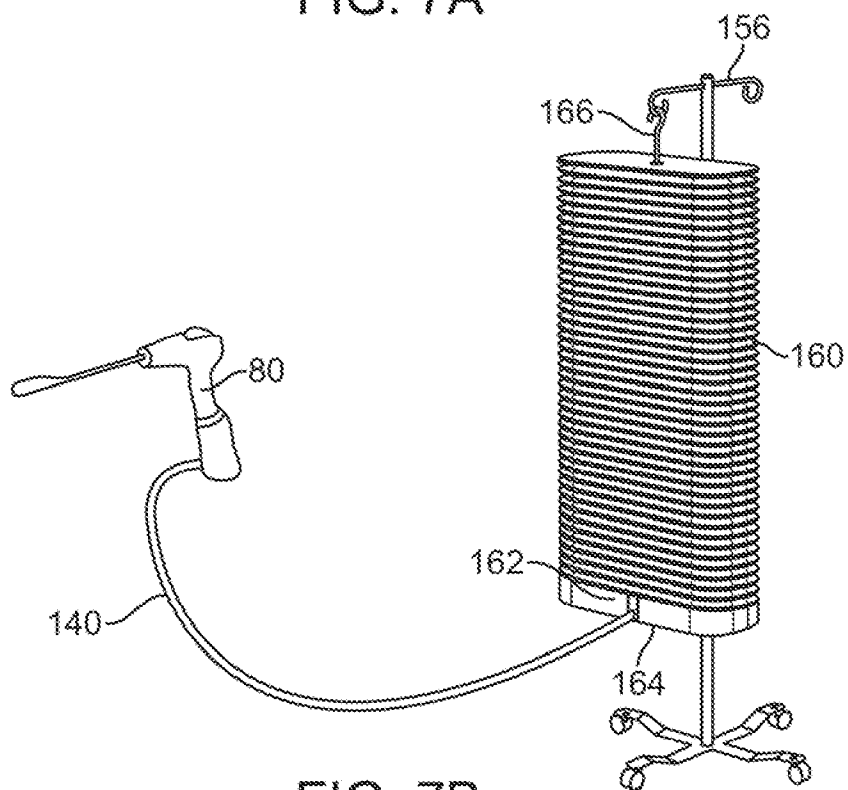


FIG. 7B

13/23

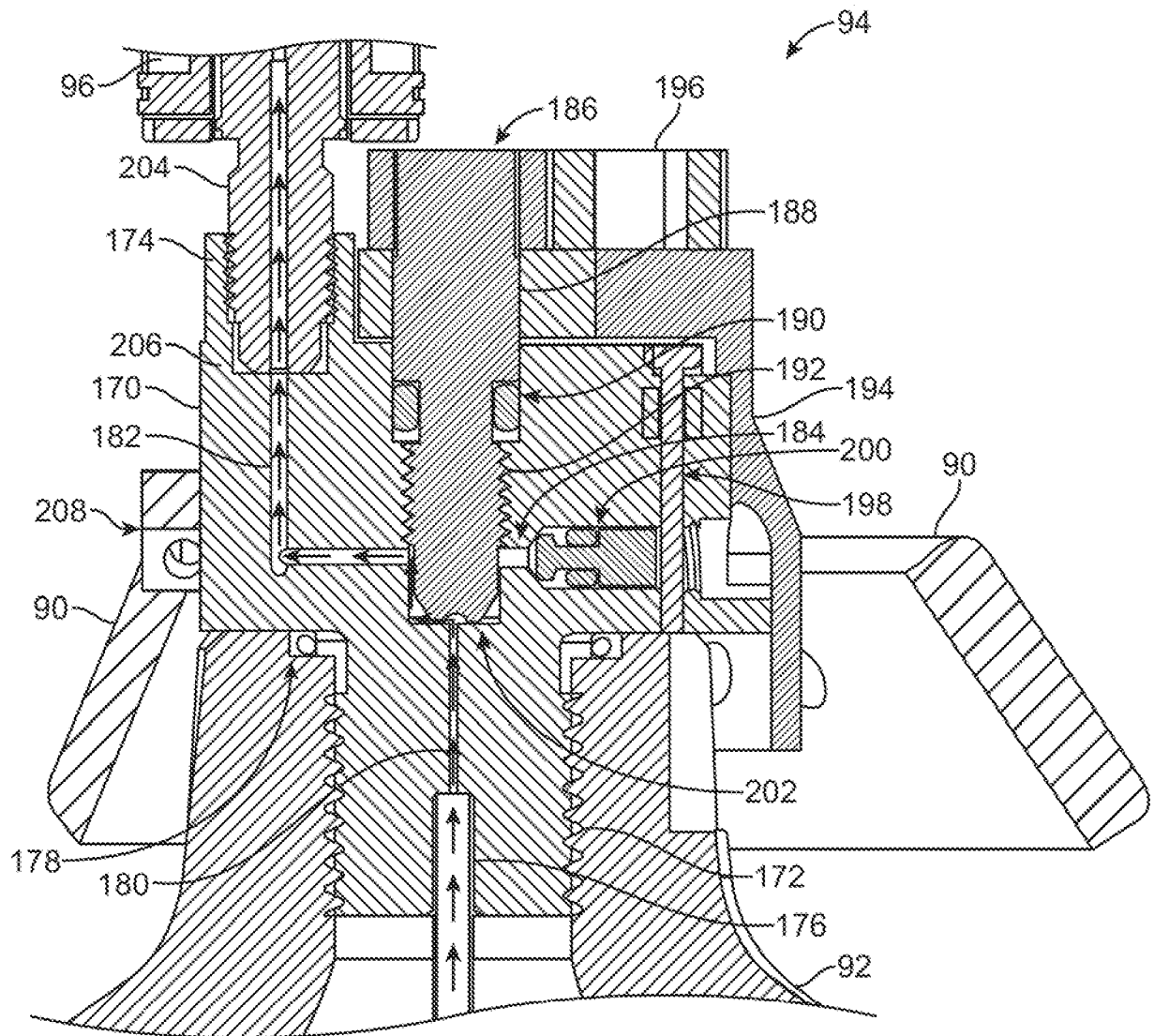


FIG. 8

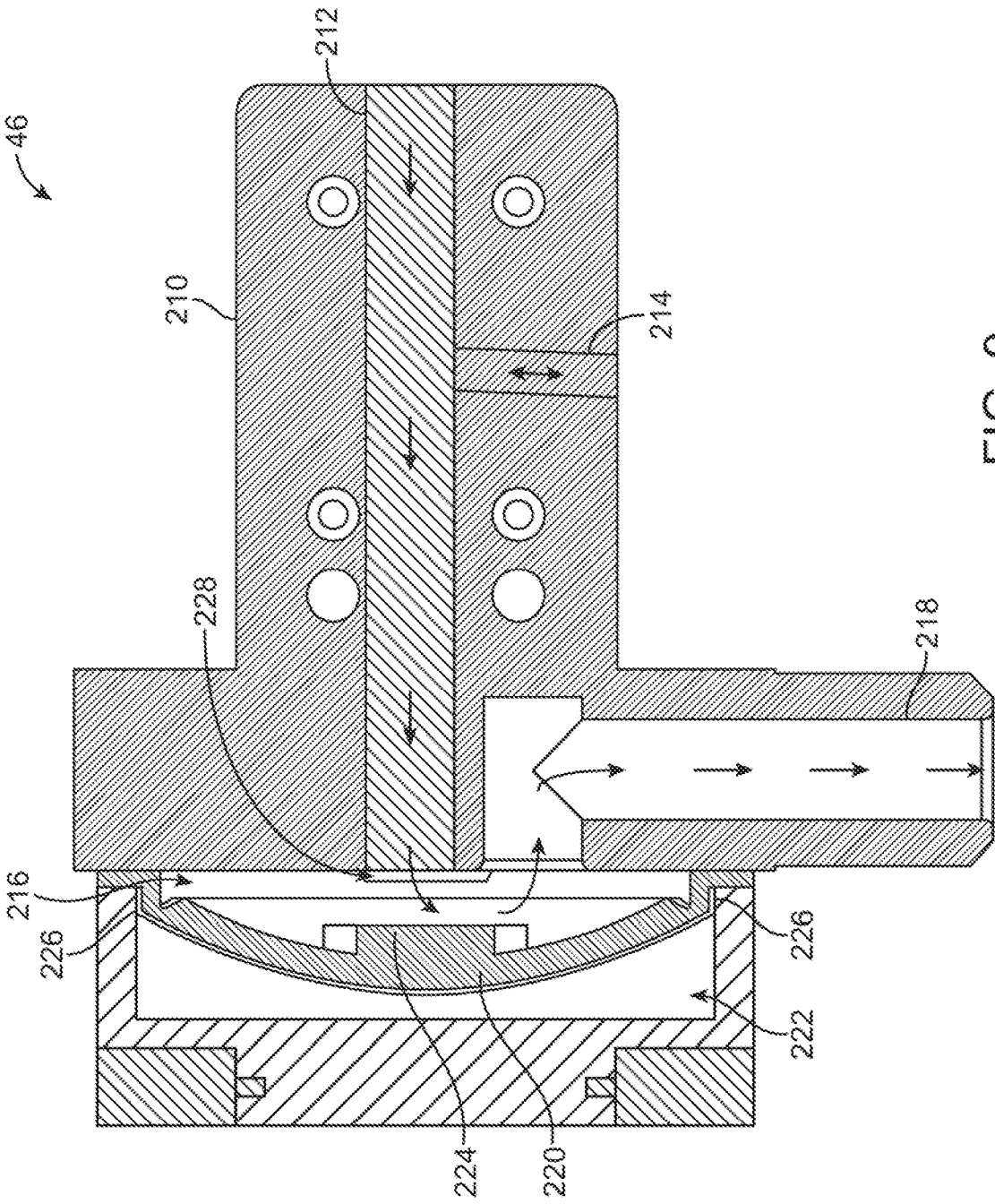


FIG. 9

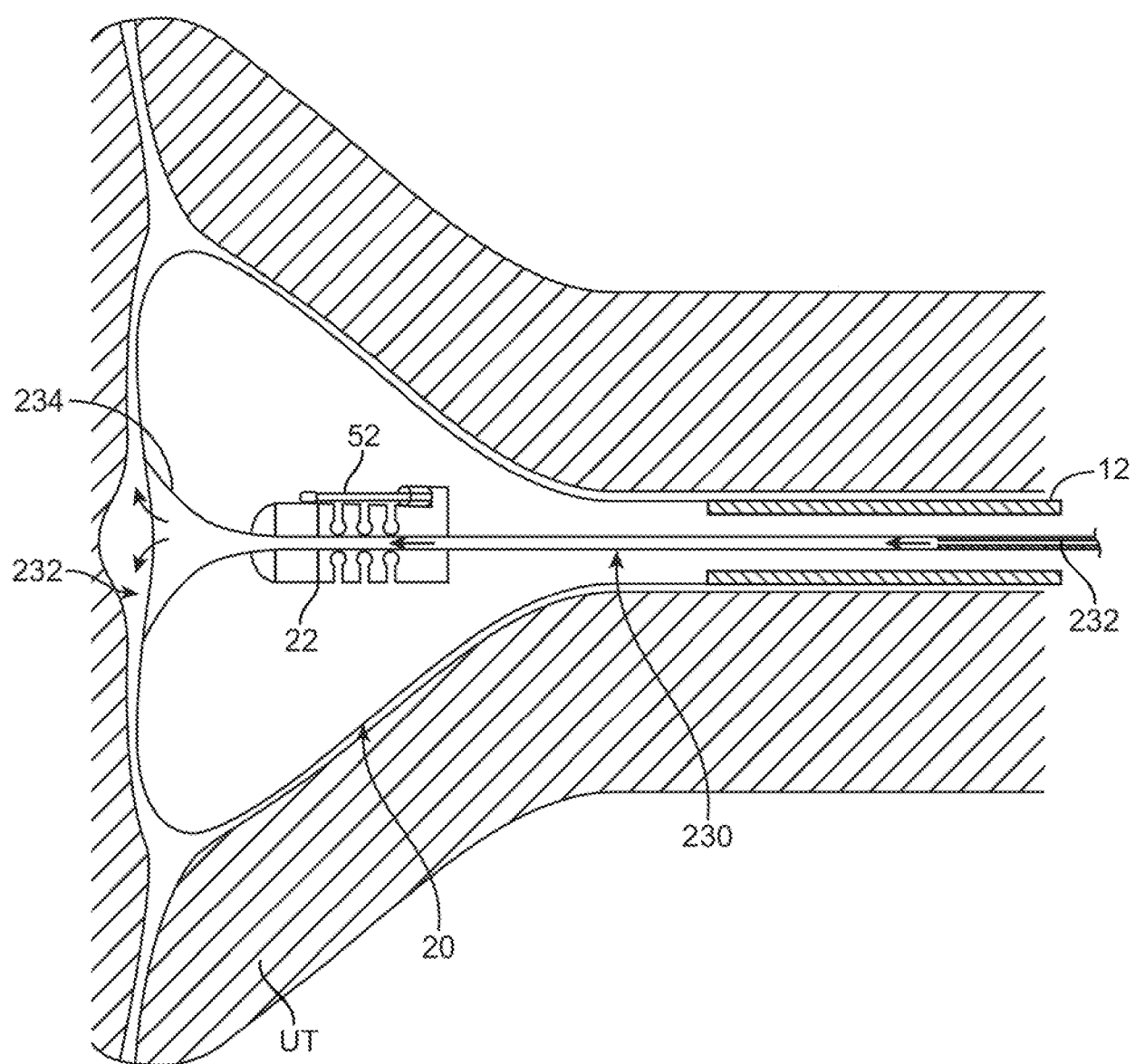


FIG. 10

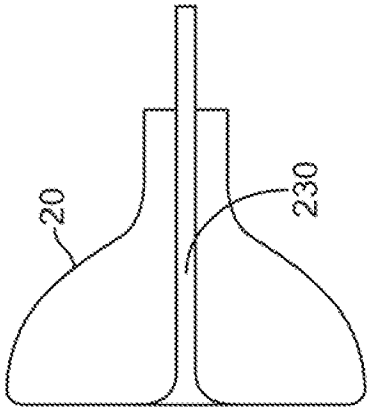


FIG. 11B

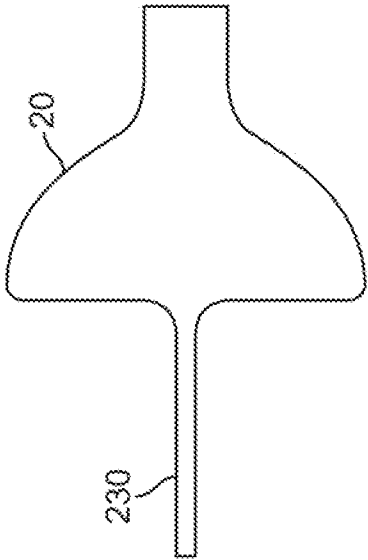


FIG. 11A

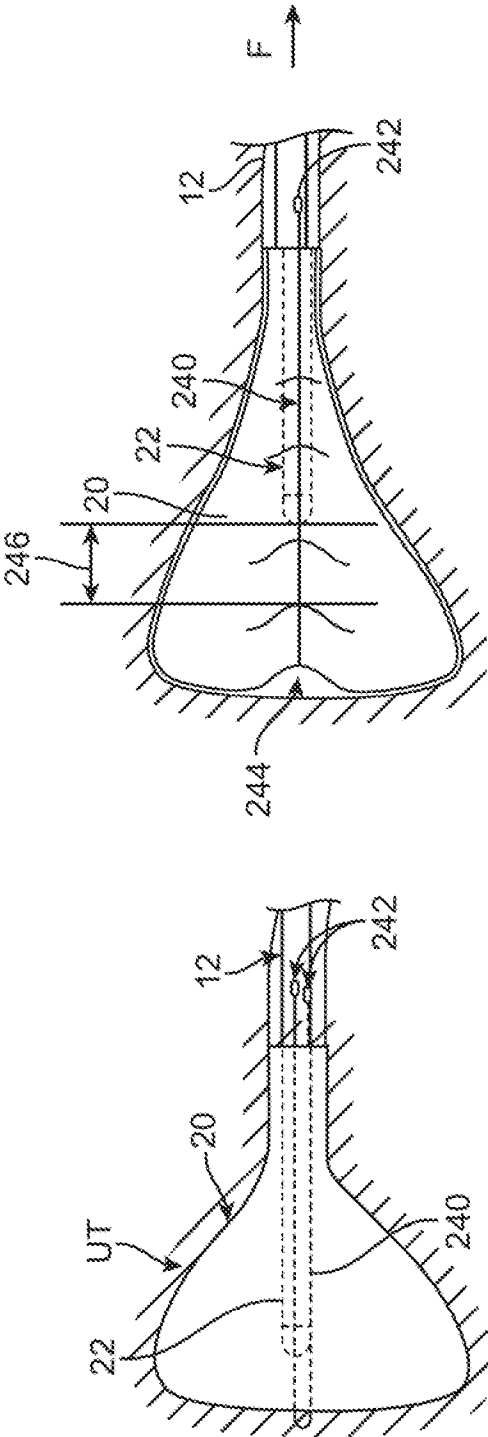


FIG. 12A

FIG. 13A

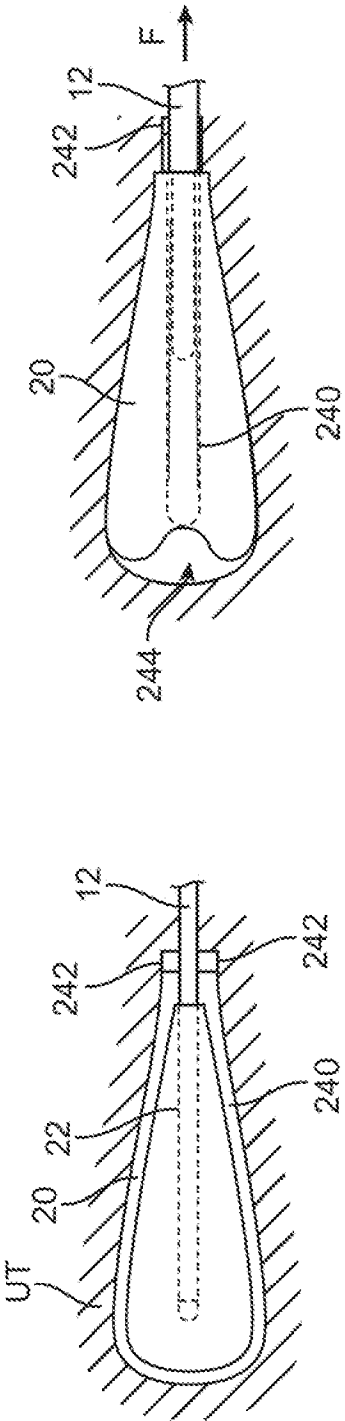


FIG. 12B

FIG. 13B

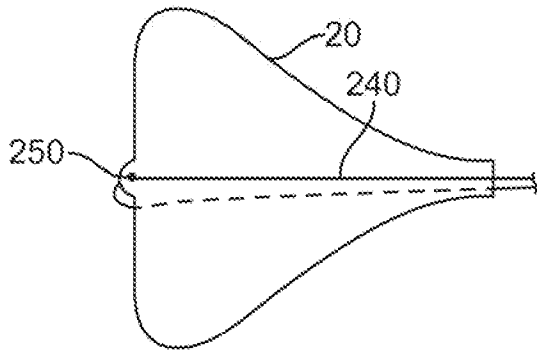


FIG. 14

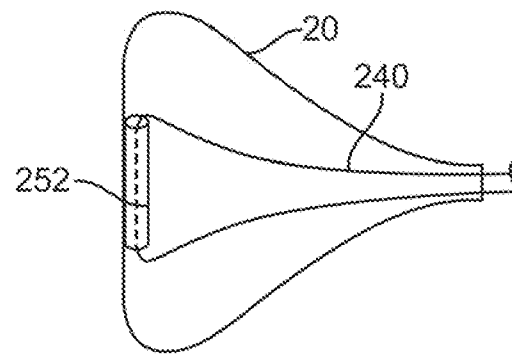


FIG. 15

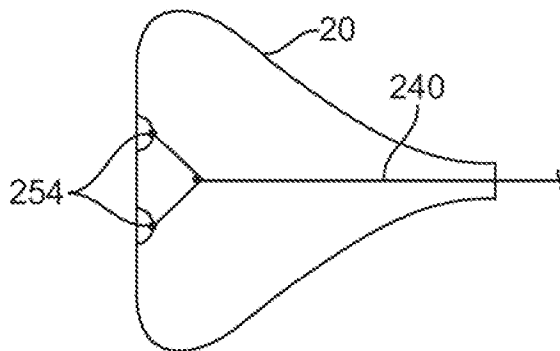


FIG. 16

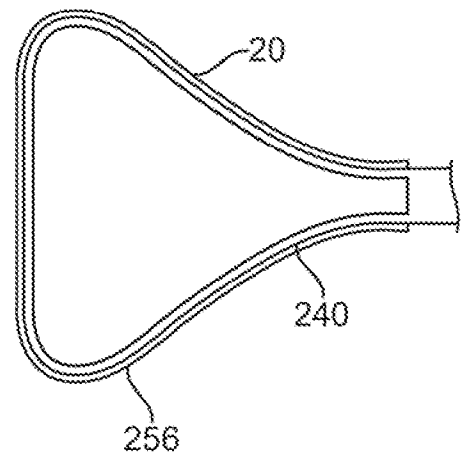


FIG. 17

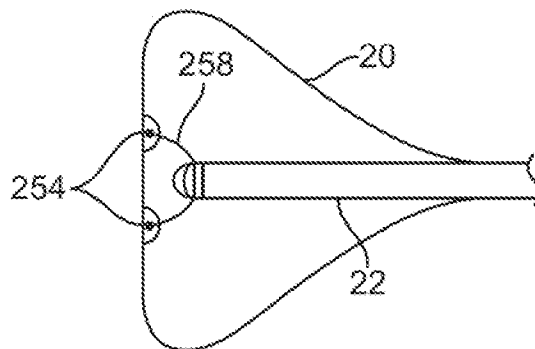


FIG. 18

19/23

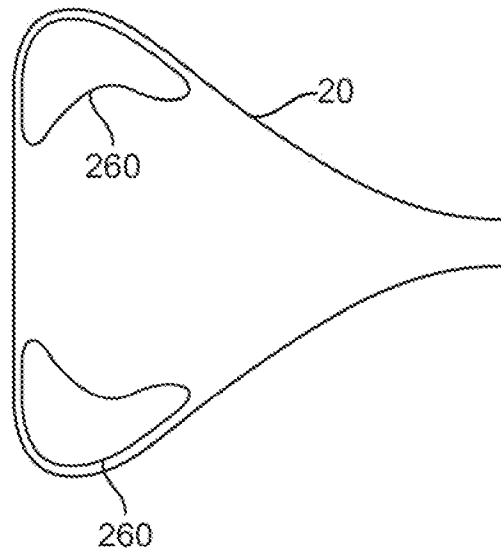


FIG. 19

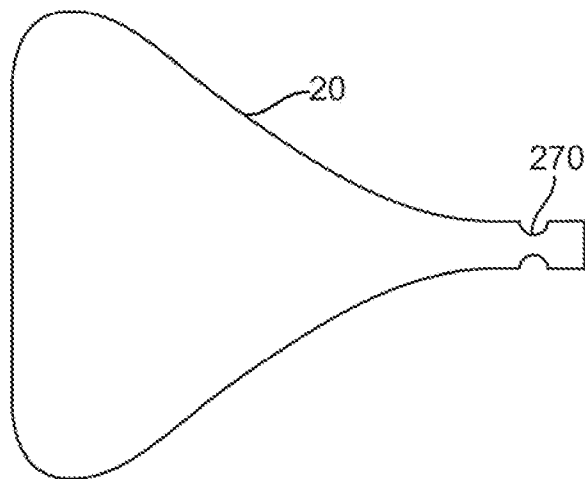


FIG. 20

20/23

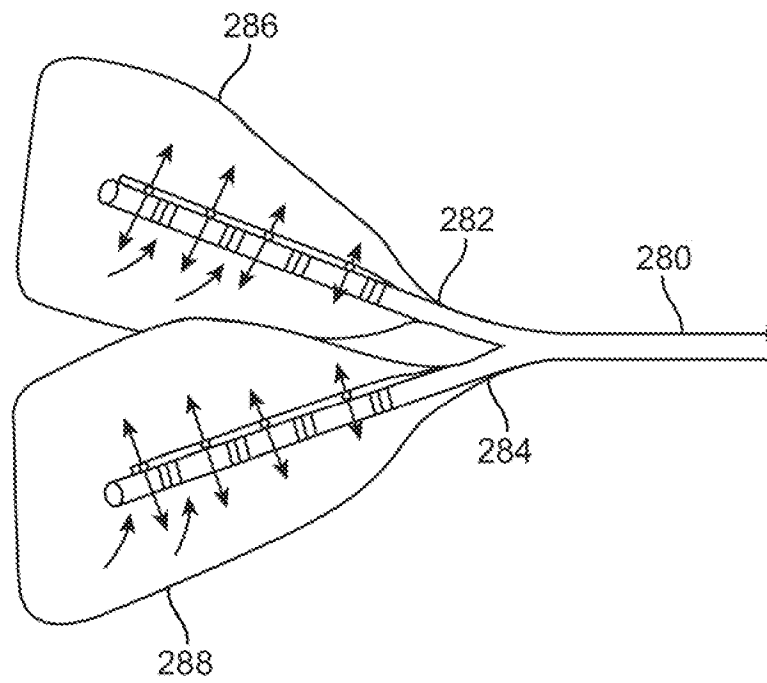


FIG. 21

21/23

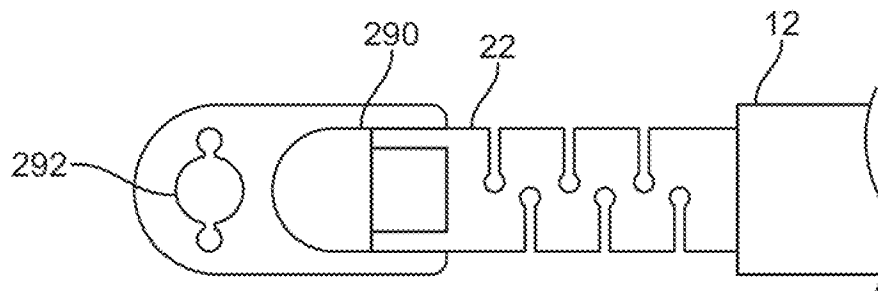


FIG. 22A

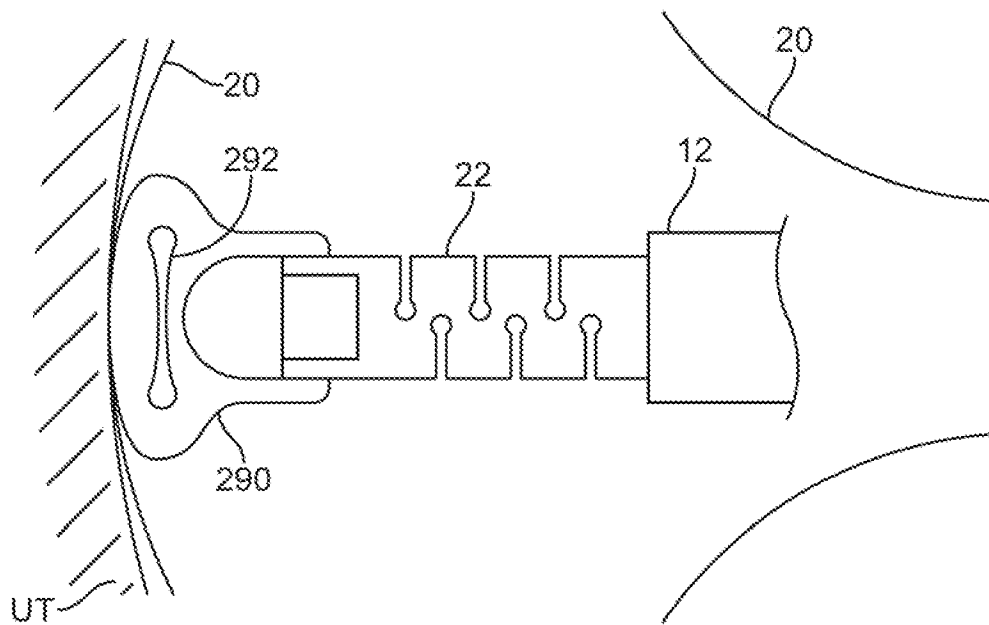


FIG. 22B

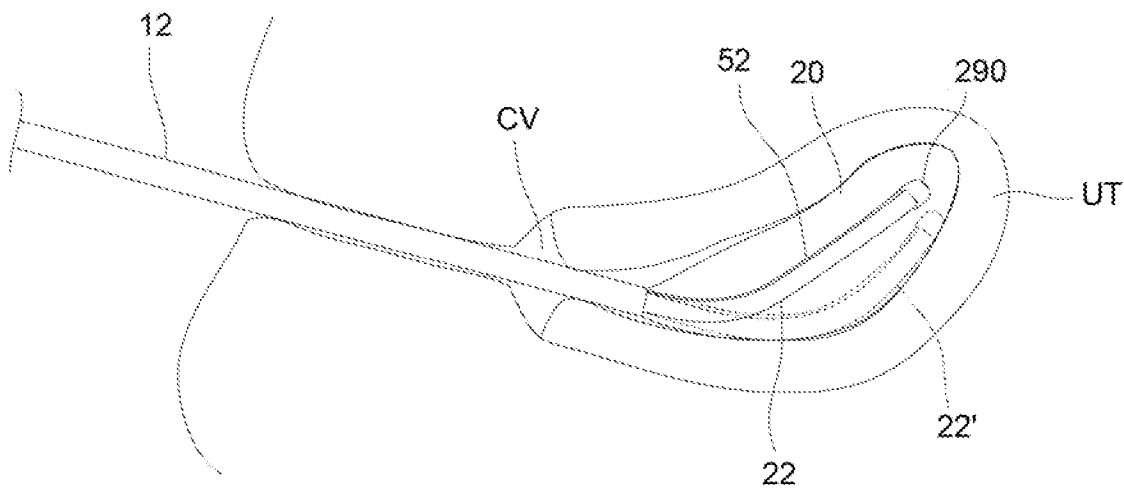
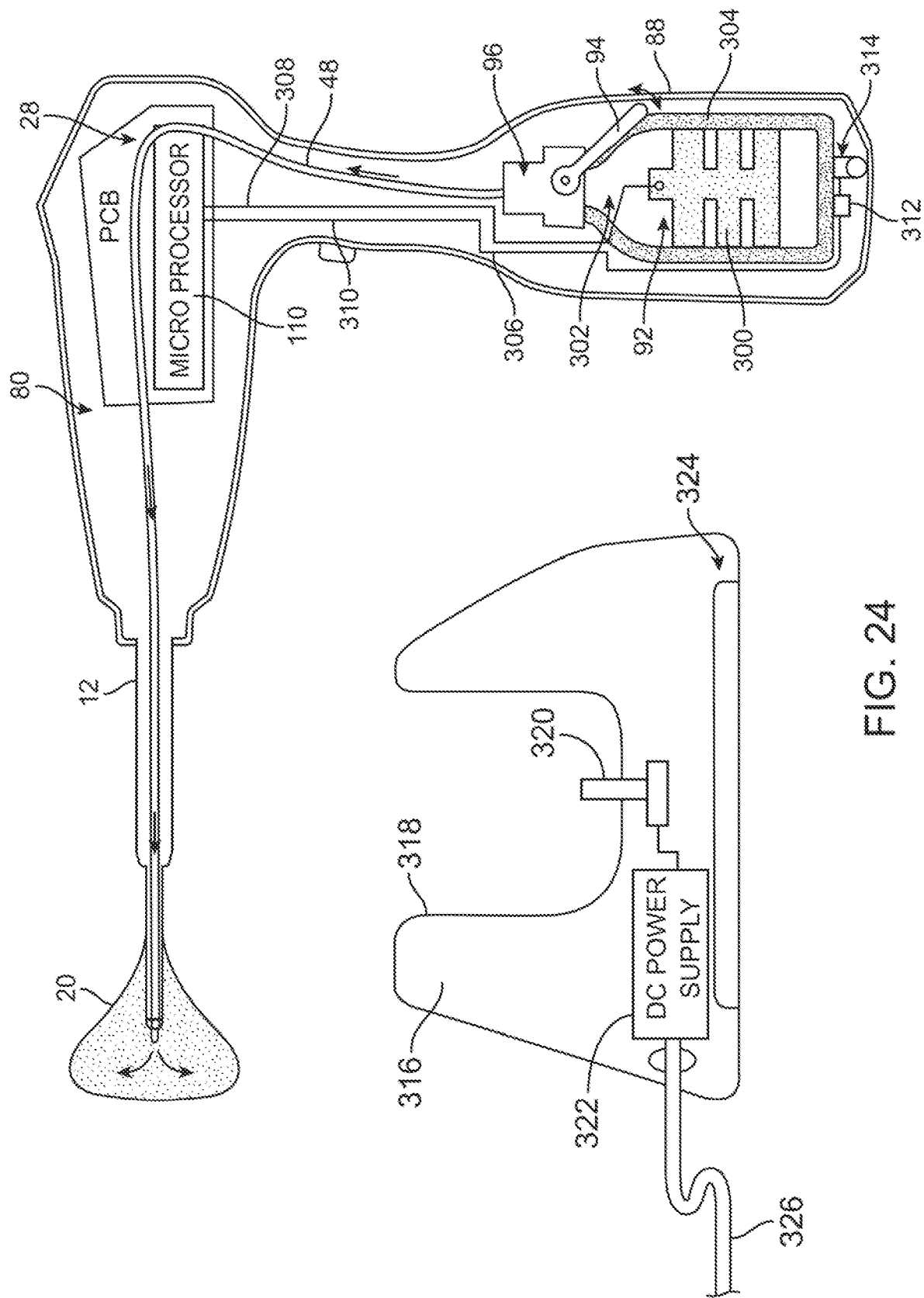


FIG. 23



INTERNATIONAL SEARCH REPORT

International application No.

PCT/US2015/017585

A. CLASSIFICATION OF SUBJECT MATTER

IPC(8) - A61B 18/02 (2015.01)

CPC - A61B 18/02 (2015.04)

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

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC(8) - A61B 17/00, 18/00, 18/02, 18/04; A61F 7/12; A61M 25/00; B25C 1/00; B65B 1/00; B67D 7/00 (2015.01)

CPC - A61B 17/00, 18/00, 18/02, 2018/0231; A61F 7/00, 7/123; A61M 25/00; B25C 1/00; B65B 1/00 (2015.04) (keyword delimited)

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

USPC - 141/82; 222/105; 227/130; 604/48, 218; 606/20, 21, 22, 28

(keyword delimited)

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

Orbit, Google Patents, Google, Google Scholar

Search terms used: canister valve, reservoir valve, container valve, cryogenic, modulated control, valve stem, inflow lumen, venting lumen, venting piston, modulated coupling, elongated probe, exhaust, liner, microprocessor

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 6,141,985 A (CLUZEAU et al) 07 November 2000 (07.11.2000) entire document	1-5, 7-10
Y		11-16, 18-21
Y	US 2012/0197245 A1 (BURNETT et al) 02 August 2012 (02.08.2012) entire document	11-16, 18-20
Y	US 2004/0167505 A1 (JOYCE et al) 26 August 2004 (26.08.2004) entire document	21
Y	US 2013/0041358 A1 (CRYOMEDIX LLC) 14 February 2013 (14.02.2013) entire document	20
A	US 3,343,544 A (DUNN et al) 26 September 1967 (26.09.1967) entire document	1-21
A	US 2009/0163902 A1 (DELONZOR et al) 25 June 2009 (25.06.2009) entire document	1-21
A	US 7,281,550 B2 (ZIEGLER) 16 October 2007 (16.10.2007) entire document	1-21

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

* Special categories of cited documents:

"A" document defining the general state of the art which is not considered to be of particular relevance

"E" earlier application or patent but published on or after the international filing date

"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)

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

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

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

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

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

"&" document member of the same patent family

Date of the actual completion of the international search

10 June 2015

Date of mailing of the international search report

01 JUL 2015

Name and mailing address of the ISA/US

Mail Stop PCT, Attn: ISA/US, Commissioner for Patents

P.O. Box 1450, Alexandria, Virginia 22313-1450

Facsimile No. 571-273-8300

Authorized officer:

Blaine R. Copenheaver

PCT Helpdesk: 571-272-4300

PCT OSP: 571-272-7774

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US2015/017585

Box No. II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)

This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☐ Claims Nos.:
because they relate to subject matter not required to be searched by this Authority, namely:

2. ☐ Claims Nos.:
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:

3. ☐ Claims Nos.:
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box No. III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:
See extra sheet.

1. ☐ As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims:
2. ☐ As all searchable claims could be searched without effort justifying additional fees, this Authority did not invite payment of additional fees.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☒ No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
1-21

Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest and, where applicable, the payment of a protest fee.
- ☐ The additional search fees were accompanied by the applicant's protest but the applicable protest fee was not paid within the time limit specified in the invitation.
- ☐ No protest accompanied the payment of additional search fees.

This application contains the following inventions or groups of inventions which are not so linked as to form a single general inventive concept under PCT Rule 13.1. In order for all inventions to be examined, the appropriate additional examination fees must be paid.

Group I, claims 1-21, are drawn to an inflow reservoir or canister valve.

Group II, claims 22-33, are drawn to a tissue treatment system.

Group III, claims 34-48, are drawn to a method of treating tissue.

The inventions listed as Groups I-III do not relate to a single general inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, they lack the same or corresponding special technical features for the following reasons:

The special technical features of Group I, an inflow reservoir or canister valve, comprising: a valve body; a reservoir interface extending from the valve body and configured for fluidly coupling with a reservoir or canister containing a cryoablative agent; a modulation control interface defined along the body and configured for fluidly coupling, to a modulation control coupler; a valve stem seated within a valve stem channel defined within the valve body; an inflow lumen defined through the valve body and extending between the reservoir interface and the modulation control interface, where the valve stem is movable between a first position which obstructs the inflow lumen and a second position which opens the inflow lumen; and a venting lumen defined through the valve body and extending between the reservoir interface and a vent opening, are not present in Groups II and III; the special technical features of Group II, a tissue treatment system, comprising: an elongate probe having a distal tip and a flexible length; at least one infusion lumen positioned through or along the elongate probe, wherein the infusion lumen defines one or more openings along its length; a liner expandably enclosing the probe; an inflow reservoir or canister valve fluidly coupled with a reservoir or canister containing a cryoablative agent; a modulation control unit fluidly coupled with the inflow reservoir or canister valve and in fluid communication with the at least one infusion lumen; and a warming element thermally coupled with the reservoir or canister, are not present in Groups I and III; and the special technical features of Group III, a method of treating tissue, comprising: monitoring a temperature or pressure of a reservoir or canister containing a cryoablative agent; maintaining the temperature or pressure of the reservoir or canister at a predetermined level; positioning an elongate probe into a body lumen to be treated; expanding a liner enclosing the probe into contact against the body lumen; and infusing a cryoablative agent through a delivery lumen such that the cryoablative agent passes into an infusion lumen, through one or more unobstructed openings, and into contact against an interior of the liner, are not present in Groups I and III.

Groups I and II share the technical feature of an inflow reservoir or canister valve fluidly coupled with a reservoir or canister containing a cryoablative agent, and a modulation control coupling with the valve. However this shared technical feature does not represent a contribution over the prior art. Specifically US 2012/0197245 A1 to Burnett et al., disclose an inflow reservoir or canister valve (coolant valve 430; FIG. 77; para. [0220]) fluidly coupled with a reservoir or canister containing a cryoablative agent (coolant reservoir 428; FIG. 77; para. [0221] "cryoablative fluid contained within the coolant reservoir 428"), and a modulation control coupling with the valve (FIG. 77; para. [0220] "coolant valve 430 which may be optionally controlled by the microcontroller 422").

Groups I and III share the technical feature of a reservoir or canister containing a cryoablative agent. However this shared technical feature does not represent a contribution over the prior art. Specifically US 2012/0197245 A1 to Burnett et al., disclose a reservoir or canister containing a cryoablative agent (coolant reservoir 428; FIG. 77; para. [0221] "cryoablative fluid contained within the coolant reservoir 428").

Groups II and III share the technical feature of a reservoir or canister containing a cryoablative agent, an elongate probe, a liner expandably enclosing the probe, and an infusion lumen having one or more openings. However this shared technical feature does not represent a contribution over the prior art. Specifically US 2012/0197245 A1 to Burnett et al., disclose a reservoir or canister containing a cryoablative agent (coolant reservoir 428; FIG. 77; para. [0221] "cryoablative fluid contained within the coolant reservoir 428"), an elongate probe (cooling probe assembly 220; FIG. 77), a liner expandably enclosing the probe (balloon 174; FIG. 77), and an infusion lumen having one or more openings (FIGS. 23A and 24 illustrate probe assembly 220 having an infusion lumen 224 having one or more openings 260; para. [0166] "cooling probe assembly 220... may comprise an exhaust catheter 222 which may define a lumen 224 therethrough" and para. [0172] "openings 260 along the catheter body 222 for venting the cryoablative fluid into the balloon 174").

Since none of the special technical features of the Group I-III inventions are found in more than one of the inventions, unity of invention is lacking.