

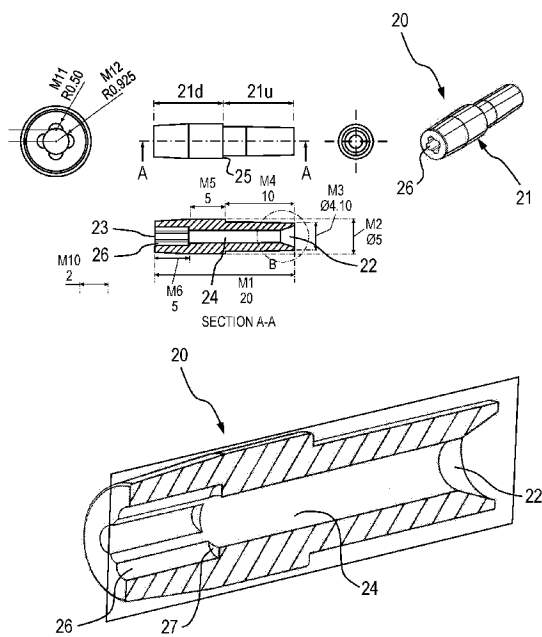


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(54) Title: LIQUID APPLICATOR



**Fig. 3**

(57) Abstract: A liquid applicator (1) for holding and discharging a curable liquid composition, comprises a receiver body (2) for holding a curable liquid composition, a discharge tip (20;30;40) having a longitudinal axis and further having a distal end remote from the receiver body (2) from which the liquid composition is discharged, and a discharge mechanism (5) for transferring liquid composition held by the applicator (1) to the tip (20;30;40) for discharge of the composition. The tip (20;30;40) comprises an outlet section (21d;31d;44d) having at least one groove formation (26;38;48) extending along the tip (20;30;40) to the distal end thereof. The applicator may be a surgical adhesive applicator.



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## Liquid Applicator

The present invention relates to a liquid applicator for holding and discharging a curable liquid composition. The invention relates particularly, but not necessarily exclusively, to such applicators intended for surgical use for successive delivery of droplets of the curable liquid composition to a surgical site. The invention has particular (but again not exclusive) application to applicators for laparoscopic surgery for the repair of hernias by fixing a mesh support material to the tissue being repaired.

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Liquid applicators which are for use in laparoscopic surgery (e.g. hernia repair operation) and which function by successive delivery of droplets (e.g. 12-15 $\mu$ l/droplet) are known in the art. An example of such an applicator is disclosed in WO 2014/072688 and WO2014/072689. A commercially available applicator (operating in accordance with the principles disclosed in the two aforementioned WO specifications) is available from Advanced Medical Solutions (Plymouth) under the name LIQUIBAND®FIX8™ and delivers droplets of a curable liquid cyanoacrylate adhesive composition. The FIX8 device is intended for use in a single surgical operation and is configured to deliver a total of about 33 droplets of adhesive, sufficient for one hernia mesh fixation operation. Cyanoacrylate is however known to interfere with natural healing at the immediate point of contact and so, for internal applications, it has been found that controlling and minimising the amount of adhesive applied is critical. To this end, liquid applicators for internal surgical use (e.g. the FIX8™ device) have outlet tips with small diameter bores (e.g. about 0.5-2.0mm). However due to the high moisture content of internal tissues there is a significant risk of the tip becoming blocked with polymerised material for one or both of two reasons. Firstly, polymerised material and tissue may build up on the outside of the tip covering the distal orifice (i.e. the orifice from which the adhesive composition is discharged onto the surgical site). Secondly, polymerised material and tissue can form inside the distal orifice causing a plug. If blockage occurs the applicator may no longer be suitable for use in completing the surgical procedure.

It is an object of the present invention to obviate or mitigate the abovementioned disadvantages.

According to the present invention there is provided a liquid applicator for holding and discharging a curable liquid composition, the applicator comprising:

5 a receiver body for holding a curable liquid composition, a discharge tip having a longitudinal axis and further having a distal end remote from the receiver body from which the liquid composition is discharged, and a discharge mechanism for transferring liquid composition held by the applicator to the tip for discharge of the composition,

wherein the tip comprises an outlet section having at least one groove formation extending along the tip to the distal end thereof.

10 The invention further provides a liquid applicator as defined in the previous paragraph charged with a curable liquid adhesive composition, e.g. contained in a frangible curable liquid adhesive composition, e.g. contained in a frangible ampoule that needs to be fractured to release the adhesive composition for discharge.

15 The liquid applicator may be configured as a surgical instrument, e.g. for use in a hernia mesh fixation operation.

We have found that the provision of at least one groove formation in an outlet section of the tip and extending therealong to the distal end thereof considerably reduces  
20 problems associated with blockage. There may be a single groove formation provided either internally or externally of the tip and having a longitudinal axis parallel to that of the tip. Preferably however a plurality of internal or external groove formations, e.g. four, are provided and have their longitudinal axes equiangularly spaced around the longitudinal axis of the tip.

25 In a particularly preferred embodiment of the invention, the tip has a bore extending through the tip to the distal end thereof and groove formations are formed in the interior wall of the bore. The bore of the tip may have an upstream section of constant cross-section (e.g. 1.5-2mm) and a downstream section in which the groove formations are  
30 provided. The groove formations may, for example, extend for a length of 4 to 6mm (preferably about 5mm).

It is particularly preferred that the upstream ends of the groove formations are defined by shoulders lying in a plane at right-angles to the direction of flow of the liquid  
35 composition through the tip.

Preferably the groove formations formed internally of the bore of the tip are of arcuate (preferably semi-circular) when seen in transverse section (i.e. the plane at right angles to the direction flow of the liquid composition through the tip).

5

In an alternative embodiment of the invention, the tip has an upstream section provided with a bore and a downstream section (providing the outlet section) in the exterior of which the groove formations are provided. In this embodiment, the grooves are each associated with an aperture providing communication between a groove and the bore (which extends no further than said apertures). The apertures may be provided at least partly in the base of each groove formation and are preferably arcuate (as seen in transverse cross-section) with each aperture extending partly up the arcuate side walls of the grooves.

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For all embodiments of the invention, it is preferred that the tip comprises a low surface energy material as such materials serve to minimise adhesion of cured adhesive composition and debris to the tip. The low surface energy material from which the tip is fabricated may, for example, comprise high density polyethylene, polypropylene, fluorinated polymer (e.g. PTFE), an acetal plastics material, silicones or a ceramic material.

20

The applicator is preferably one adapted to deliver successive droplets of curable liquid composition, e.g. by virtue of a trigger mechanism configured to effect the successive discharge. The droplets may have a volume of 10-20 $\mu$ l, (e.g. 12-15 $\mu$ l) and the device may be configured for delivery of 25-40, e.g. 30-35, droplets of adhesive.

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The applicator may comprise an elongate cannula through which the adhesive composition is discharged and at the end of which the tip is mounted. For this purpose, the tip may comprise an upstream body portion and a downstream body portion of lower cross-sectional size than the former. The upstream body portion may therefore be in the form of a spigot for location in the end of the cannula with a shoulder (formed at the junction of the upstream and downstream body portions) abutting against the end of the cannula. The upstream and downstream body portions may for example be of circular cross-section. In preferred embodiments of the invention, the cannula incorporates an inner tube (e.g. of a fluorinated polymer such a PTFE) along which the

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adhesive composition is fed from the body of the applicator to the tip. This inner tube preferably extends into the tip and preferably terminates before the outlet end thereof.

5 The applicator is preferably one configured for laparoscopic surgical use. It will however be appreciated that applicators in accordance with the invention may be used for open surgery.

10 In the applicator configured for surgical use, the tip may, for example, have an overall length of 15-25mm (e.g. 18-24mm) and/or a maximum cross-sectional dimension of 3-8mm (e.g. 4-6mm). If sub-divided into upstream and downstream body portions, the subdivision may be about midway along the length of the tip. The bore that extends into the tip from the upstream end thereof (and which feeds the adhesive composition to the groove formations) may optionally have a converging inlet section (for insertion of the aforementioned inner tube) but otherwise be of uniform, cross-section to the point where it communicates with the upstream ends of the groove formations. Other cross-sections may however be employed, e.g. the bore may narrow in cross-section part-way along its length at a converging step formation before it reaches the upstream ends of the groove formations. Apart from a converging inlet section, the bore may, for example, have a maximum cross-sectional size of 1.5 to 3.5mm.

20 In the embodiments as described above, the groove formations may, for example, have a length of 4-8mm. In the case where the groove formations have an arcuate (e.g. semi-circular section) then the radius of that section may, for example, be 0.3 to 1mm (e.g. 0.4 to 0.8mm). As viewed in transverse cross-section of the tip (i.e. in a plane intersected at right angles by the longitudinal axis of the tip) the centres of the arcuate sections may be 0.75mm to 1.5mm from the longitudinal axis of the tip. The or each groove formation preferably has a longitudinal axis that extends parallel to the longitudinal axis of the tip.

30 A liquid adhesive applicator in accordance with the invention intended for surgical use is ideally such that there is no substantial polymerisation (curing) of the adhesive before it reaches the end of the tip so that curing occurs when the adhesive comes into contact with the moisture of tissue being repaired. As such, it is preferred that there are no polymerisation initiators pre-deposited on the tip upstream of the outlet end

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thereof (such as are provided in certain other types of applicators of curable adhesive compositions).

The curable liquid composition is preferably a cyanoacrylate adhesive composition.

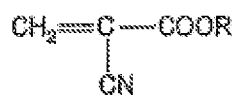
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Examples of cyanoacrylate adhesive compositions that may be incorporated in the applicator are given below.

10 The adhesive fluids that may be applied by the applicator of the present invention may be comprised of a wide variety of cyanoacrylate adhesive formulations. The reservoir may contain a stronger bonding and less flexible cyanoacrylate adhesive composition, such as *n*-butyl cyanoacrylate, or it may contain a more flexible tissue adhesive, such as an octyl or hexyl or decyl or other homologs of cyanoacrylate.

15 Preferably, the cyanoacrylate compositions used comprise cyanoacrylate prepolymer compositions that can be applied as a liquid/gel to the skin surface. Optionally, the cyanoacrylate prepolymers can include therapeutic agents such as analgesics, anti-inflammatory agents, antimicrobial agents, and the like.

20 Preferably, the polymerizable cyanoacrylate prepolymers comprise cyanoacrylate esters that, in monomeric form, are represented by the formula I:



I

wherein

25

R is selected from the group consisting of:

alkyl of 1 to 10 carbon atoms,

alkenyl of 2 to 10 carbon atoms,

cycloalkyl groups of from 5 to 8 carbon atoms,

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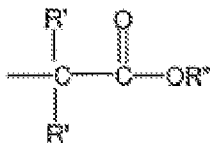
phenyl,

2-ethoxyethyl,

3-methoxybutyl,

6

and a substituent of the formula:



7

wherein

5

each R' is independently selected from the group consisting of:

hydrogen and methyl, and

R'' is selected from the group consisting of:

alkyl of from 1 to 6 carbon atoms,

10

alkenyl of from 2 to 6 carbon atoms,

alkynyl of from 2 to 6 carbon atoms,

cycloalkyl of from 3 to 8 carbon atoms,

aralkyl selected from the group consisting of benzyl, methylbenzyl and phenylethyl,

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phenyl, and phenyl substituted with 1 to 3 substituents selected from the group consisting of hydroxyl, chloro, bromo, nitro, alkyl of 1 to 4 carbon atoms, and alkoxy of from 1 to 4 carbon atoms.

20 More preferably, in the cyanoacrylate esters of formula I, R is an alkyl group of from 2 to 10 carbon atom including ethyl, *n*-propyl, iso-propyl, *n*-butyl, iso-butyl, sec-butyl, *n*-pentyl, iso-pentyl, *n*-hexyl, 2-ethylhexyl, *n*-heptyl, octyl, nonyl, and decyl. Mixtures of such compounds can also be employed as disclosed by Berger, et al., U.S. Patent No. 5,998,472, which is incorporated herein by reference in its entirety.

25 It is to be understood that the term "polymerizable cyanoacrylate esters" refers to polymerizable formulations comprising cyanoacrylate monomers or polymerizable oligomers which, in their monomeric form, are preferably compounds represented by formula I as described above.

The polymerizable cyanoacrylate esters described herein rapidly polymerize in the presence of water vapour or tissue protein, and the *n*-butyl-cyanoacrylate bonds to mammalian skin tissue without causing histotoxicity or cytotoxicity.

- 5 Polymerizable cyanoacrylate esters are known in the art and are described in, for example, U.S. Patent Nos. 3,527,224; 3,591,676; 3,667,472; 3,995,641; 4,035,334; and 4,650,826 the disclosures of each are incorporated herein by reference in their entirety.
- 10 Optionally, the cyanoacrylate compositions applied by the present applicator can include a "biocompatible plasticizer". As used herein, the "biocompatible plasticizer" refers to any material which is soluble or dispersible in the cyanoacrylate composition, which increases the flexibility of the resulting polymeric film coating on the skin surface, and which, in the amounts employed, is compatible with the skin as measured by the
- 15 lack of moderate to severe skin irritation. Suitable plasticizers are well known in the art and include those disclosed in U.S. Patent Nos. 2,784,127 and 4,444,933 the disclosures of both of which are incorporated herein by reference in their entirety. Specific plasticizers include, by way of example only, acetyl tri-*n*-butyl citrate (preferably ~20 weight percent or less), acetyl trihexyl citrate (preferably ~20 weight
- 20 percent or less) butyl benzyl phthalate, dibutyl phthalate, dioctylphthalate, *n*-butyryl tri-*n*-hexyl citrate, diethylene glycol dibenzoate (preferably ~20 weight percent or less) and the like. The particular biocompatible plasticizer employed is not critical and preferred plasticizers include dioctylphthalate and C<sub>2</sub>-C<sub>4</sub>-acyl tri-*n*-hexyl citrates.
- 25 Optionally as well, the cyanoacrylate composition applied by the present applicator can include an "antimicrobial agent". As used herein, the term "antimicrobial agent" refers to agents which destroy microbes (i.e. bacteria, fungi, yeasts and viruses) thereby preventing their development and their pathogenic action.
- 30 Preferred cyanoacrylate compositions useful in the practice of this invention are also disclosed by Greff, et al., U.S. Patent No. 5,480,935, which application is incorporated herein by reference in its entirety. In a particularly preferred embodiment, the cyanoacrylate adhesive composition further comprises an antimicrobially effective amount of compatible antimicrobial agent. Such compositions preferably comprise
- 35 from 0.1 to about 30 and preferably about 0.5 to 10 weight percent of the compatible

antimicrobial agent either as a solution or as suspension based on the total weight of the composition. Compatible antimicrobial agents are those which are either soluble or suspendable in the cyanoacrylate composition, which do not cause premature polymerization of the cyanoacrylate composition, which do not prevent polymerization of the cyanoacrylate composition when applied to mammalian skin, and which are compatible with the intended use including biocompatibility with the patient's skin. Suitable such compositions are disclosed in U.S. Patent No. 6,475,502 , which discloses compositions of cyanoacrylate/povidone-iodine complexes, and US 2005-0042196 A1, which discloses compositions of cyanoacrylate esters and phenol. All three disclosures are incorporated herein by reference in their entirety.

The use of compatible antimicrobial agent in the compositions permits the agent to be released from the polymeric film thereby reducing microbial growth adjacent to the film.

Other medicaments suitable for use in conjunction with the cyanoacrylate compositions include corticoid steroids such as described by Greff, et al. in U.S. Patent No. 5,962,010 which is incorporated herein by reference in its entirety and analgesic compounds such as lidocaine. The former reduces inflammation whereas the latter reduces pain. Combinations of a steroid with an analgesic are also covered.

The invention will be further described, by way of example only, with reference to the accompanying drawings, in which:

Fig 1 shows a commercially available surgical adhesive applicator;

Fig 2 shows views of a conventional applicator tip as supplied with the applicator shown in Fig 1;

Fig 3 shows views of a first embodiment of applicator tip for use in accordance with the invention.

Fig 4 shows views of a second embodiment of applicator tip for use in accordance with the invention; and

Fig 5 shows views of a third embodiment of applicator tip for use in accordance with the invention.

Illustrated in Fig 1 is an adhesive applicator 1 sold by Advanced Medical Solutions (Plymouth) Ltd under the trade mark "LIQUIBAND<sup>®</sup>FIX8<sup>™</sup>" for use in fixing a support mesh (usually of polypropylene) in place during a hernia repair operation, particularly

such an operation effected laparoscopically. The “LIQUIBAND<sup>®</sup>FIX8<sup>™</sup>” device is intended for use in a single hernia repair operation during which a plurality of drops of curable cyanoacrylate adhesive composition are selectively and successively discharged to fix the mesh in position. The illustrated applicator 1 is of the type disclosed in WO 2014/072688 and WO 2014/072689 the disclosure of which is hereby incorporated by reference and to which reference may be made for full details as to the manner in which the applicator 1 is constructed and operated. However, in brief, applicator 1 comprises a body unit 2, an elongate cannula 3 extending from the body unit 2 and provided at its free distal end with an applicator tip 4, and further provided with a trigger mechanism 5. Although not illustrated in Fig 1, cannula 3 has a liner tube of a fluorinated polymer which extends along cannula 3 from its proximal end within body unit 2 into the bore of the applicator tip 4 to a position level with the free distal end of tip 4. As supplied, the FIX8<sup>™</sup> device includes in the body unit 2 thereof a frangible ampoule (not shown) containing a liquid adhesive composition incorporating a curable cyanoacrylate adhesive composition which is curable on contact with patient tissue to which the aforementioned support mesh is to be adhered. Provided on and within the body unit 2 are mechanisms which allow the ampoule to be fractured to release the adhesive and then to allow the cannula 3 (or more specifically the fluorinated liner tube) to be primed more-or-less along its full length with adhesive so that operation of the trigger mechanism 5 allows (for each such operation) discharge from the tip 4 of a droplet of liquid adhesive of a fixed volume (about 10-15 $\mu$ l). The amount of adhesive supplied with the FIX8<sup>™</sup> applicator is sufficient for discharge of about 33 droplets of adhesive.

During a hernia repair operation, the aforementioned mesh is located in position against the patient's internal tissue to be repaired and droplets of adhesive (discharged as briefly described above by operation of trigger mechanism 5) are applied to the junctions of the mesh (on the side remote from patient tissue) so that adhesive can flow over the junction and into contact with the tissue (to form an adhesive “anchor”) and cure, whereby the mesh becomes bonded to the tissue.

The tip 4 currently supplied with the FIX8<sup>™</sup> device is shown in Fig 2 and will be seen to comprise a tubular body 11 with an inwardly converging inlet 12 and an outlet 13 connected by a bore 14 which is of uniform, circular cross-section along its length from the downstream end of converging inlet 12 to the outlet 13. The provision of the

inwardly converging inlet 12 facilitates insertion of the aforementioned fluorinated liner tube into the tip 4.

5 Body 11 is provided approximately mid-way along its length with an external annular shoulder 15 which sub-divides body 11 into upstream and downstream sections 11u and 11d respectively. Upstream section 11u is for location within the distal end of cannula 3 such that annular shoulder 15 abuts against the distal end of cannula 3 with the downstream end 11d projecting beyond the end of cannula 3.

10 Tip 4 as supplied with the FIX8™ device has the dimensions shown in Table 1 below.

**Table 1**

Length of body 11	20mm
Length of upstream section 11u	10mm
Length of downstream section 11d	10mm
Outside diameter of upstream section 11u	5mm
Height of shoulder 15	0.9mm
Diameter of bore 14	1.85mm

15 As indicated in Table 1, bore 14 has an internal diameter of about 1.85mm. This allows small quantities of adhesive to be applied (about 13µl with each activation of the trigger mechanism 5), which is advantageous since cyanoacrylate is known to interfere with natural healing at the immediate point of contact with patient tissue. Therefore, for internal applications, it has been found that controlling and minimising the amount of  
 20 adhesive applied is critical. However, due to the high moisture content of internal tissues, coupled with the narrow diameter of bore 14, issues arise with the interior of the tip becoming blocked with polymerised cyanoacrylate material (and also polymerised cyanoacrylate around the exterior of the distal end of the tip 4). If the blockage cannot be cleared then the applicator cannot be used for further application of  
 25 adhesive.

Reference is now made to Fig. 3 which illustrates an embodiment of tip 20 which is for use in an applicator in accordance with the invention and which has been found significantly to reduce problems associated with blockage from polymerised

cyanoacrylate. Thus, for example, tip 20 may be used in the FIX8™ device in place of the tip 4 described with reference to Fig. 2. The illustrated tip 20 has some general similarity with tip 4 and as such comprises a generally tubular body 21 having an inwardly converging inlet 22 and an outlet 23 connected by a bore 24 that extends from the downstream end of inwardly converging inlet 22 to the outlet 23. Body 21 is provided with an external annular shoulder 25 which, in effect, sub-divides body 21 into upstream and downstream sections 21u and 21d whereof the former is of lower external diameter than the latter. Upstream body section 21u locates in the distal end of cannula 3 with the shoulder 25 locating against the distal end of cannula 3.

Formed in the wall of the bore 24 are four axially parallel, equiangularly spaced grooves 26 that extend from a location about halfway along downstream body section 21d to the outlet 23. More specifically, the grooves 26 extend in the downstream direction (to the outlet 23) from respective upstream end faces 27 formed as a result of the grooves 26 being moulded in the walls of block 24. Grooves 26 are generally semi-circular as seen in transverse cross-section (i.e. in a plane at right angles to the longitudinal axis of tip 20) and their configuration is best seen in the cut-away sectional view (which is to a much enlarged scale) in the lowermost drawing of Fig. 3.

Tip 20 may have the dimensions shown in Table 2 below.

**Table 2**

Length of body 21	20mm
Length of upstream section 21u	10mm
Length of downstream section 21d	10mm
Outside diameter of upstream section 21u	5mm
Height of shoulder 25	0.9mm
Diameter of bore 24	1.85mm
Lengths of grooves 26	5mm
Radius of grooves 26	0.5mm with centre positioned 0.8mm from centre of distal tip

Figure 4 illustrates a further embodiment of tip 30 that may be used in an applicator in accordance with the invention to alleviate the above described blockage problems. Tip

30 has some similarity in overall construction with tip 20 and therefore, for convenience parts in tip 30 that have a counterpart in tip 20 are designated by a reference with the same final numeral and ten more than the corresponding numeral in Fig. 3. Thus, for example, the body of tip 30 is referenced as 31 (cf body 21 of tip 20). Also, for convenience, description of the like parts is not repeated.

In contrast with the embodiment of applicator tip 20 illustrated in Fig. 3, the applicator tip 30 is provided with external grooves 38 extending parallel to the longitudinal axis of tip 30 and being open along their length at the outer surface of tip 30. Each groove 38 is generally arcuate in transverse cross-section and extends between the distal end 33 of tip 30 and a respective wall 39 at the upstream end of groove 38. Over their lengths, the upper edges of grooves 38 rise slightly in going from distal end 33 to wall 39 whereby the grooves 38 increase in depth in going in this direction. Formed in the base of each groove 38 at the upstream end thereof (and extending part way up the lateral surfaces of the grooves) are respective apertures 39 that provide communication with the bore 34, which in this embodiment extends from the downstream end of inwardly converging inlet section 34 to the downstream ends of apertures 38. Thus, when tip 30 is located in the end of cannula 3 of applicator 1 and the applicator is primed with adhesive, operation of trigger mechanism 5 causes adhesive to pass into bore 34 of tip 30 and then *via* apertures 39 into the grooves 38 for discharge from tip 30.

Tip 30 may have the dimensions shown in Table 3 below.

**Table 3**

Length of body 31	20mm
Length of upstream section 31u	10mm
Length of downstream section 31d	10mm
Outside diameter of upstream section 31u	5mm
Height of shoulder 35	0.9mm
Diameter of bore 34	1.85mm
Lengths of grooves 38	5mm

Radius of grooves 38	0.75mm radius with the centre positioned 1.3mm from the centre of the distal tip
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A further embodiment of tip 40 is illustrated in Fig. 5 and, like tip 30 shown in Fig. 4, is provided with external grooves referred as 48, although in this case the grooves 48 are of constant depth along their length is going from distal end 43 of tip 40 to end face 49.

5 Tip 40 includes a stepped bore shown as having an upstream section 44u and narrower downstream section 44d which extends a short distance beyond the upstream ends of grooves 48. Provided on the base of each groove 48 of the upstream end is a respective aperture 39 providing communication between a groove 38 and the downstream section 44d of the bore.

10

Tip 4 may have the dimensions shown in Table 4 below.

**Table 4**

Length of body 41	23mm
Diameter of upstream bore section 44u	3.2mm
Diameter of downstream bore section 44d	4.53mm
Lengths of grooves 48	7.68mm
Radius of grooves 48	0.6mm with the centre positioned 1.35mm from the centre of the distal tip

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To demonstrate the effectiveness of the invention, samples of tips as shown in Figs. 2-5 and having the dimensions indicated in the respective Tables above were manufactured from PTFE and further samples were manufacture from a Class VI acetal plastics material (Delrin®). The various tips were tested for their blockage characteristics using a test rig reliably simulating the delivery characteristics of the FIX8 device (repeated delivery of *ca* 13µl droplets of a curable cyanoacrylate adhesive composition as supplied with the device). In all tests, the adhesive composition was delivered to the tips through a fluorinated polymer tube of the type used in the FIX8

20

device. The tube was inserted into the upstream end of the tip and extended to a position 5mm from the discharge end of the tip.

5 The tests involved bonding a polypropylene mesh (of the type used in a hernia fixation operation) to chicken tissue using the various tips at angles of 90° to the horizontal (i.e. tip pointing vertically downwards), 60° to the horizontal (tip inclined downwardly), and 75° to the vertical (tip inclined upwards).

For each tip at each angle, the test was conducted using the following procedure.

10

(A) An adhesive anchor (provided by a droplet of adhesive) was expressed on to the mesh/chicken tissue surface.

15

(B) After each anchor application, the mesh was pushed and held in place using the tip for 10 seconds.

(C) After a wait of 20 seconds, the next adhesive anchor was deployed.

20

(D) The rate of adhesive delivery was such that at least 5 anchors were delivered within 2 minutes.

(E) The tip was observed for blockage due to debris accumulation after each adhesive anchor delivery.

25

(F) After deployment of every 3 adhesive anchors, the tip was subjected to the moist surface on the chicken tissue for 10 seconds.

(G) Steps (A)-(F) were repeated until at least 33 anchors had been expressed onto the mesh/chicken sample or until tip blockage (whichever occurred earlier).

30

As a result of the tests, it was found that tips of the type shown in Fig. 2 and made from either PTFE or acetal plastics material were prone to blockage that could not be cleared before 33 anchors had been expressed.

The tip shown in Fig. 3 and made of acetal plastics material provided the best results of all tips tested and was able to deliver adhesive composition for the required minimum of 33 anchors even though debris had accumulated around the tip. In the case of the tip of Fig. 3 made from PTFE, accumulated debris blocked the tip but the block was clearly visible and could easily be removed to restore the flow of adhesive to allow expression of the minimum 33 anchors.

In the case of the tips shown in Fig. 4 and 5 (for both PTFE and acetal plastics) it was found that the adhesive bleed holes did not block completely but debris did accumulate on the four external grooves. The debris could easily be removed to allow continued flow of adhesive but it was observed there was some difficulty in detecting whether or not adhesive reached the target site due to the accumulation of debris at the tip.

**CLAIMS:**

1. A liquid applicator for holding and discharging a curable liquid composition, the applicator comprising:
  - 5 a receiver body for holding a curable liquid composition, a discharge tip having a longitudinal axis and further having a distal end remote from the receiver body from which the liquid composition is discharged, and a discharge mechanism for transferring liquid composition held by the applicator to the tip for discharge of the composition, wherein the tip comprises an outlet section having at least one groove formation  
10 extending along the tip to the distal end thereof.
2. An applicator as claimed in claim 1 wherein said at least one groove formation has a longitudinal axis parallel to the longitudinal axis of the tip.
- 15 3. An applicator as claimed in claim 2 wherein a plurality of said groove formations are provided.
4. An applicator as claimed in claim 3 wherein four of said groove formations are provided.  
20
5. An applicator as claimed in claim 3 or 4 wherein said groove formations have their longitudinal axis equiangularly spaced around the longitudinal axis of the tip.
6. An applicator as claimed in claim 4 or 5 wherein the tip has a bore extending  
25 through the tip to the distal end thereof and the groove formations are formed in the interior wall of the bore.
7. An applicator as claimed in claim 6 wherein the bore has an upstream section of constant cross-section and a downstream section in which said groove formations are  
30 provided.
8. An applicator as claimed in claim 6 or 7 wherein the groove formations are arcuate in transverse cross-section.

9. An applicator as claimed in claim 8 wherein the groove formations are semi-circular in transverse cross-section.
10. An applicator as claimed in any one of claims 6 to 9 wherein the upstream ends of groove formations are defined by shoulders lying in a plane at right angles to the direction of flow of the liquid composition through the tip.
11. An applicator as claimed in any one of claims 3 to 5 wherein said tip has an upstream section provided with a bore and a downstream section providing the outlet section and in the exterior of which the groove formations are provided, said groove formations each being associated with an aperture providing communication between a groove and the bore.
12. An applicator as claimed in claim 11 wherein an aperture is provided at least partly in the base of each of the groove formations.
13. An applicator as claimed in claim 12 wherein the groove formations are arcuate as seen in cross-section transverse to the direction of flow of liquid adhesive composition and each aperture provided extends partly up the arcuate side walls of the groove formations.
14. An applicator as claimed in any one of claims 1 to 13 wherein the tip comprises a low surface energy material.
15. An applicator as claimed in claim 14 wherein the low surface energy material is a fluorinated polymer, e.g. PTFE.
16. An applicator as claimed in claim 14 wherein the low surface energy material is an acetal plastics material.
17. An applicator as claimed in any one of claims 1 to 16 adapted to deliver successive droplets of curable liquid composition.
18. An applicator as claimed in claim 17 wherein the droplets have a volume of 10-20 $\mu$ l, e.g. 12-15 $\mu$ l.

19. An applicator as claimed in claim 17 or 18 configured for delivery of 25-40 droplets of adhesive composition.

5 20. An applicator as claimed in any one of claims 17 to 19 comprising a trigger mechanism configured to effect said successive discharge of droplets.

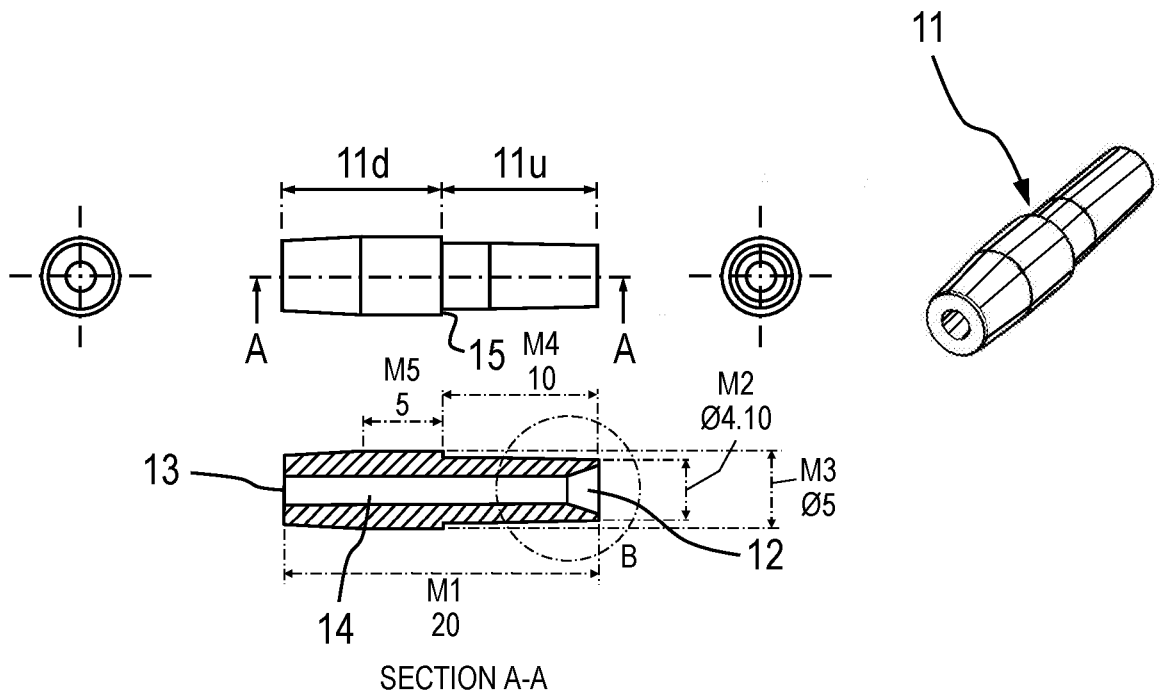
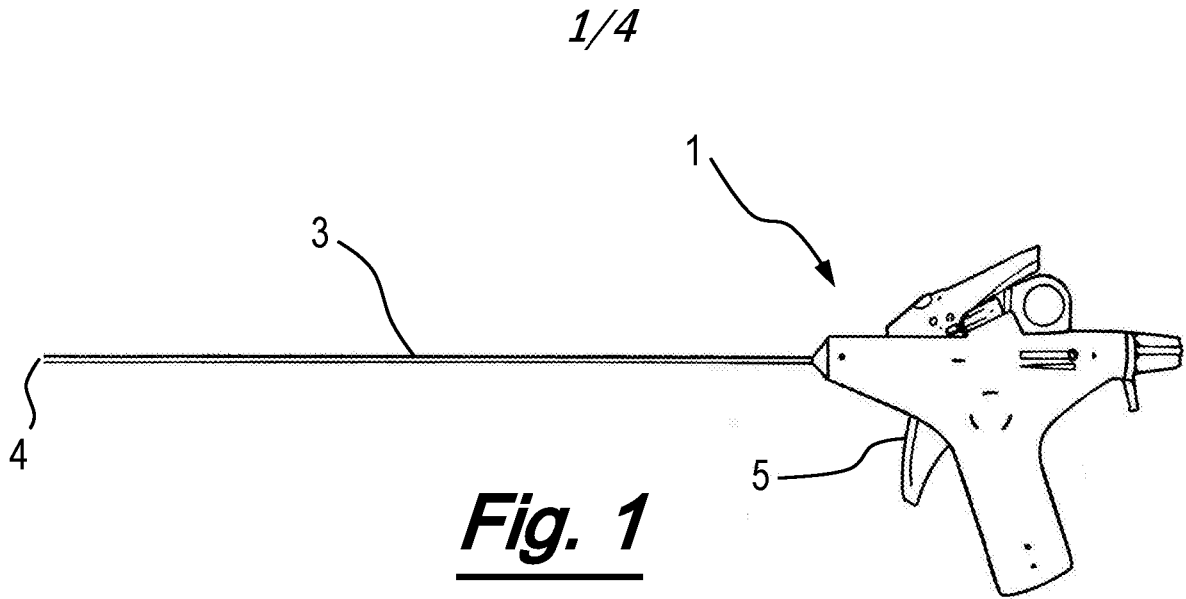
10 21. An applicator as claimed in any one of claims 1 to 20 having an elongate cannula through which the adhesive composition is discharged and at the end of which the tip is mounted.

22. An applicator as claimed in any one of claims 1 to 21 for use in laparoscopic surgery.

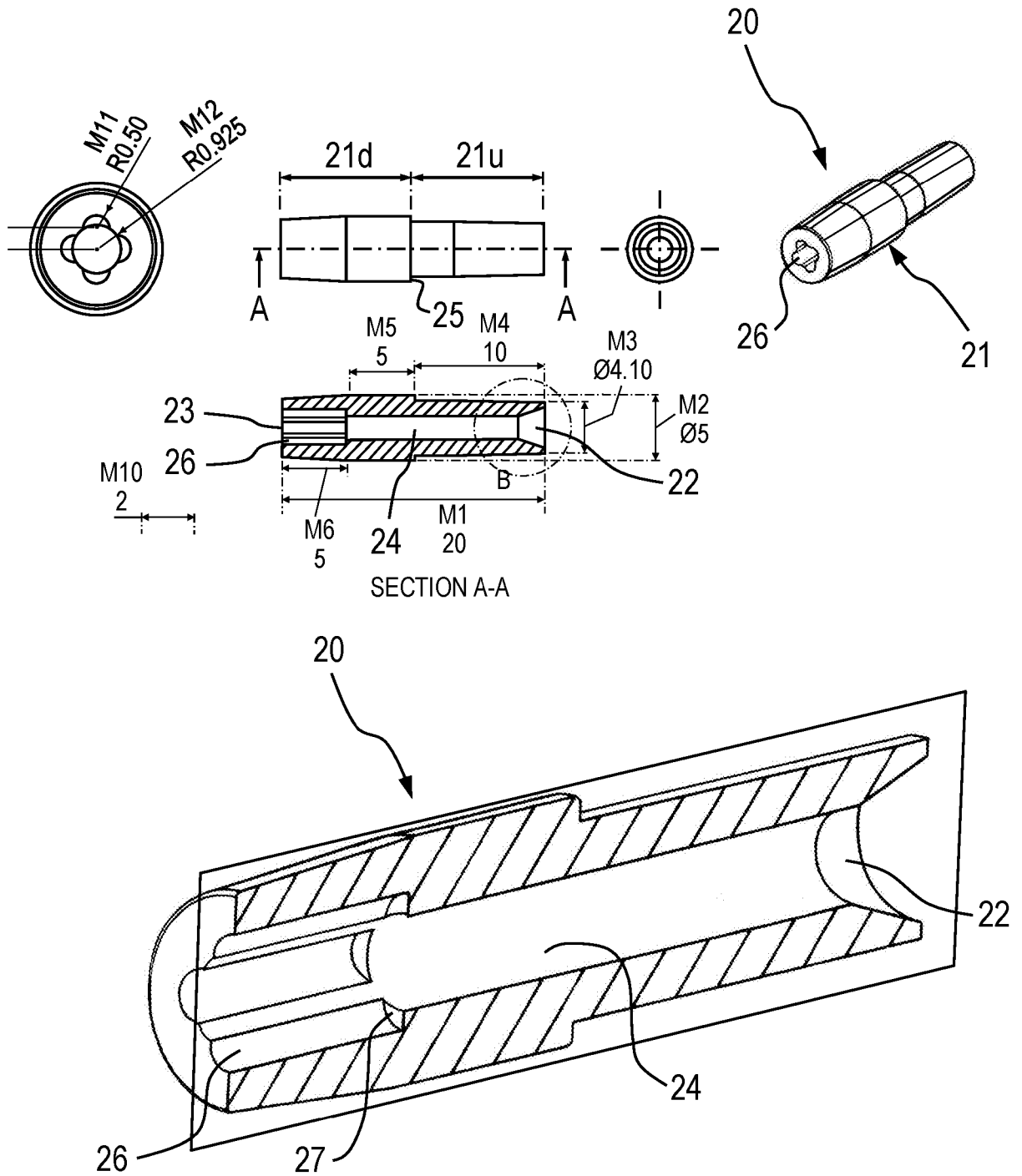
15 23. An applicator as claimed in any one of claims 1 to 22 charged with curable adhesive composition.

20 24. An applicator as claimed in claim 23 wherein the curable liquid composition is a cyanoacrylate adhesive composition.

25 25. An applicator as claimed in claim 23 wherein the curable liquid composition comprises n-butyl cyanoacrylate.



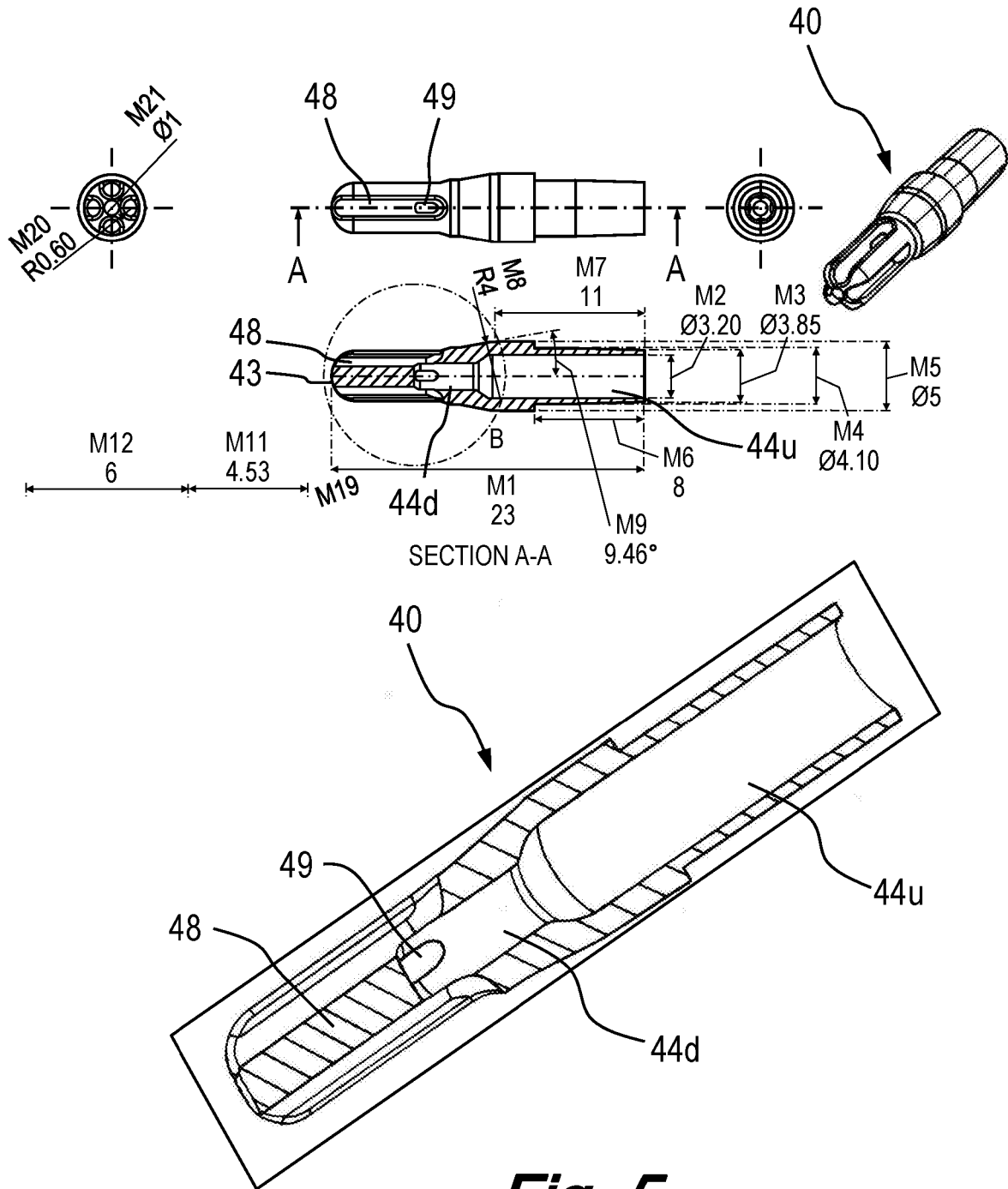
**Fig. 2**



**Fig. 3**



4/4



**Fig. 5**

INTERNATIONAL SEARCH REPORT

International application No  
PCT/GB2017/051260

A. CLASSIFICATION OF SUBJECT MATTER  
INV. A61B17/00 B05B1/06  
ADD.  
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)  
A61B B05B

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

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)  
EPO-Internal, WPI Data

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 2002/176733 A1 (CLARK JEFFREY G [US] ET AL) 28 November 2002 (2002-11-28)	1-3, 11-13, 23-25
Y	paragraphs [0014], [0016], [0044] - [0050], [0057], [0072], [0082], [0086]; figures 4-8	8,9
X	CH 709 684 A1 (MEDMIX SYSTEMS AG [CH]) 30 November 2015 (2015-11-30) paragraphs [0036] - [0037], [0055] - [0059], [0064]; figures 1, 13-15	1-5,11, 12,21-23
X	US 2013/072984 A1 (ROBINSON JAMES C [US]) 21 March 2013 (2013-03-21)	1-5,11, 12,17-22
Y	paragraph [0041] - paragraph [0046]; figures 7A, 8-11, 13,	8,9
	----- -/--	

Further documents are listed in the continuation of Box C.  See patent family annex.

\* Special categories of cited documents :

<p>"A" document defining the general state of the art which is not considered to be of particular relevance</p> <p>"E" earlier application or patent but published on or after the international filing date</p> <p>"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)</p> <p>"O" document referring to an oral disclosure, use, exhibition or other means</p> <p>"P" document published prior to the international filing date but later than the priority date claimed</p>	<p>"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</p> <p>"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</p> <p>"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</p> <p>"&amp;" document member of the same patent family</p>
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Date of the actual completion of the international search  18 July 2017	Date of mailing of the international search report  28/07/2017
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Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Fax: (+31-70) 340-3016	Authorized officer  Kink, Thomas
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## INTERNATIONAL SEARCH REPORT

International application No  
PCT/GB2017/051260

C(Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 2015/041559 A1 (ALBISETTI NICOLAS [FR]) 12 February 2015 (2015-02-12)	1-7,10, 17-20
Y	paragraphs [0065], [0102], [0112]; figures 7, 8	8,9
X	WO 2010/097896 A1 (AQUA SCIENCE CORP [JP]; HAYASHIDA ATSUSHI [JP]) 2 September 2010 (2010-09-02)	1-7, 17-20
Y	paragraphs [0006] - [0008], [0019] - [0023]; figures 1, 3, 4	8,9
X	EP 2 638 969 A2 (NORDSON CORP [US]) 18 September 2013 (2013-09-18)	1-6, 17-20
Y	paragraphs [0047], [0059]; figures 1,7	8,9
X	JP 2011 020031 A (TERUMO CORP) 3 February 2011 (2011-02-03)	1,14,15
Y	paragraphs [0026], [0037] - [0039], [0050], [0061], [0066], [0071] - [0072]; figures 1-4	16
X	US 2001/004692 A1 (KIDOOKA SATOSHI [JP] ET AL) 21 June 2001 (2001-06-21)	1,21,22
Y	paragraphs [0001], [0022], [0024] - [0026]; figures 1-3	
Y	GB 2 462 136 A (MEDLOGIC GLOBAL LTD [GB]) 3 February 2010 (2010-02-03)	16
A	page 3, line 14 - line 25 page 8, line 12 - line 24 page 9, line 16 - line 24 page 11, line 9 - line 22 page 14, line 1 - line 6 page 18, line 17 - line 20 figures 1-6, 8, 11	1-15, 17-25
A	WO 2014/071395 A2 (SMITH & NEPHEW INC [US]) 8 May 2014 (2014-05-08)	1-4,17, 18,20
Y	paragraph [0042] - paragraph [0043]; figure 1	
A	US 2009/112255 A1 (LEOPOLD PHILLIP M [US] ET AL) 30 April 2009 (2009-04-30)	17-19
Y	claims 1, 4; figure 4	

# INTERNATIONAL SEARCH REPORT

Information on patent family members

International application No PCT/GB2017/051260
---

Patent document cited in search report	Publication date	Patent family member(s)	Publication date	
US 2002176733	A1	28-11-2002	US 6428233 B1 US 2002176733 A1	06-08-2002 28-11-2002
CH 709684	A1	30-11-2015	CH 709684 A1 CN 106535778 A EP 3145417 A1 US 2017100115 A1 WO 2015176905 A1	30-11-2015 22-03-2017 29-03-2017 13-04-2017 26-11-2015
US 2013072984	A1	21-03-2013	US 2013072984 A1 US 2013072986 A1 WO 2013043763 A1	21-03-2013 21-03-2013 28-03-2013
US 2015041559	A1	12-02-2015	CN 104144751 A EP 2800633 A1 FR 2985202 A1 RU 2014126620 A US 2015041559 A1 WO 2013102866 A1	12-11-2014 12-11-2014 05-07-2013 20-02-2016 12-02-2015 11-07-2013
WO 2010097896	A1	02-09-2010	NONE	
EP 2638969	A2	18-09-2013	CA 2808957 A1 CN 103301556 A EP 2638969 A2 JP 2013188479 A US 2013245576 A1	13-09-2013 18-09-2013 18-09-2013 26-09-2013 19-09-2013
JP 2011020031	A	03-02-2011	JP 5588131 B2 JP 2011020031 A	10-09-2014 03-02-2011
US 2001004692	A1	21-06-2001	JP 4441025 B2 JP 2001169997 A US 2001004692 A1	31-03-2010 26-06-2001 21-06-2001
GB 2462136	A	03-02-2010	NONE	
WO 2014071395	A2	08-05-2014	AU 2013337240 A1 CA 2889609 A1 CN 104883984 A EP 2914181 A2 JP 2015534852 A US 2015265821 A1 WO 2014071395 A2	14-05-2015 08-05-2014 02-09-2015 09-09-2015 07-12-2015 24-09-2015 08-05-2014
US 2009112255	A1	30-04-2009	AU 2008317048 A1 BR PI0818771 A2 CA 2703337 A1 CN 101883597 A EP 2211954 A2 JP 2011502006 A KR 20100086006 A RU 2010121159 A US 2009112255 A1 WO 2009055322 A2	30-04-2009 07-04-2015 30-04-2009 10-11-2010 04-08-2010 20-01-2011 29-07-2010 10-12-2011 30-04-2009 30-04-2009

专利名称(译)	液体涂抹器		
公开(公告)号	<a href="#">EP3451936A1</a>	公开(公告)日	2019-03-13
申请号	EP2017723486	申请日	2017-05-05
[标]申请(专利权)人(译)	高级药液		
申请(专利权)人(译)	ADVANCED MEDICAL SOLUTIONS LIMITED		
当前申请(专利权)人(译)	ADVANCED MEDICAL SOLUTIONS LIMITED		
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发明人	MILLER, GUY STEPHEN GOPALAKRISHNAN, NITHINKRISHNAN PARISH, SIMON MARK		
IPC分类号	A61B17/00 B05B1/06		
CPC分类号	A61B17/00491 A61B2017/005 A61B2017/00522 A61B2017/0084 A61B2017/00853 A61B2090/037 A61L24/06		
优先权	2016007868 2016-05-05 GB		
外部链接	<a href="#">Espacenet</a>		

#### 摘要(译)

一种用于保持和排出可固化液体组合物的液体涂布器 ( 1 ) , 包括用于保持可固化液体组合物的接收器主体 ( 2 ) , 具有纵向轴线并且还具有远端远程的排出尖端 ( 20; 30; 40 ) 从用于排出液体组合物的接收器主体 ( 2 ) 和用于将由涂敷器 ( 1 ) 保持的液体组合物转移到尖端 ( 20; 30; 40 ) 以排出组合物的排出机构 ( 5 ) 。尖端 ( 20; 30; 40 ) 包括出口部分 ( 21d; 31d; 44d ) , 其具有至少一个沿尖端 ( 20; 30; 40 ) 延伸到其远端的凹槽结构 ( 26; 38; 48 ) 。涂抹器可以是外科粘合剂涂抹器。