



(22) Date de dépôt/Filing Date: 1991/12/31

(41) Mise à la disp. pub./Open to Public Insp.: 1992/07/27

(45) Date de délivrance/Issue Date: 2003/02/18

(30) Priorités/Priorities: 1991/01/26 (P 41 02 336.6) DE;
1991/11/14 (91 119 406.6) DE

(51) Cl.Int.⁵/Int.Cl.⁵ G01N 35/00, G01N 1/10

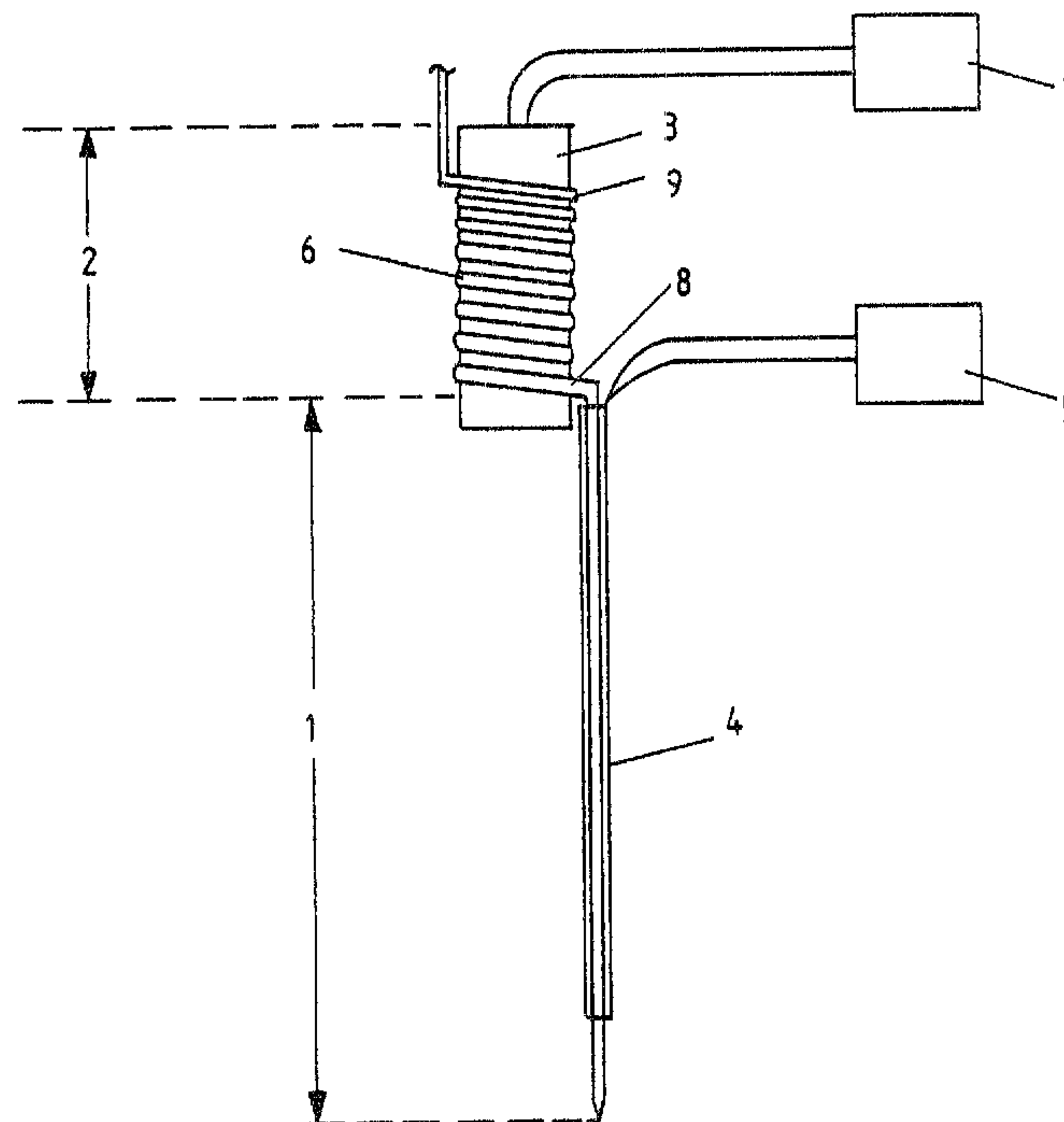
(72) Inventeurs/Inventors:
GEIB, MANFRED, DE;
HEIDE, WOLFGANG, DE;
WITTEKIND, JURGEN, DE;
BREETZ, MICHAEL, DE

(73) Propriétaire/Owner:
DADE BEHRING MARBURG GMBH, DE

(74) Agent: BERESKIN & PARR

(54) Titre : PIPETTE

(54) Title: PIPETTE TUBE



(57) Abrégé/Abstract:

A heatable pipette apparatus, which includes a pipette tube having a first zone and second zone. The first zone includes a tip end and a first heater for heating the first zone. The second zone includes a second heater for heating the second zone. Regulators are provided for separately regulating the temperature of the first heater and the second heater, the first heater being able to more accurately and precisely alter the temperature of fluids in the tube to reach a predetermined temperature.



ABSTRACT OF THE DISCLOSURE

A heatable pipette apparatus, which includes a pipette tube having a
5 first zone and second zone. The first zone includes a tip end and a first heater
for heating the first zone. The second zone includes a second heater for
heating the second zone. Regulators are provided for separately regulating
the temperature of the first heater and the second heater, the first heater
being able to more accurately and precisely alter the temperature of fluids in
10 the tube to reach a predetermined temperature.

TITLE: PIPETTE TUBE**BACKGROUND OF THE INVENTION****Field of the Invention**

5 The invention relates to a heatable pipette tube with a controlled electric heater.

Description of the Related Art

10 A heatable pipette tube, such as the one described in German patent specification 26 26 332, permits regulatable heating extending over the pipette tube, with the exception of the tip end. A drawback of this design is that temperature control of samples is difficult. For example, if the heating and the regulation (control) is adjusted in order to temper larger volumes, there is a danger that smaller volumes may become overheated. Likewise, if the
15 pipette is adjusted for small volumes, large samples may not become heated enough. There also exists a controllable heater for a centrifuge, into which a pipette tube may be inserted. However, temperature control with this arrangement is also problematic. That is, it is difficult to temper different volumes of pipette fluid quickly to a desired temperature without overheating
20 the fluid.

 EP-0 192 957 B1 describes an analysis device with a slide which transports via a pipette, fluid samples from a rotating plate to a tempered cuvette. The slide is provided with a controllable heater. This temperature controller pretempers the sample fluid to the temperature of the cuvette.

25 German Offenlegungsschrift 38 38 626 discloses a pipette tube that is heated by electrically heated water. However, this heater lacks sufficient responsiveness because the water supply together with other structural parts must first be heated by the water to the desired temperature.

SUMMARY OF THE INVENTION

An aspect of the present invention is to provide a heatable pipette tube in which different volumes can be brought very quickly to the desired temperature.

5 This aspect is attained by providing a pipette tube with two zones, namely a first zone containing the tip of the pipette tube and comprising a relatively small pipette volume, and a second zone with a comparatively larger pipette volume. Both zones are provided with separate heaters each with its own regulator. The regulator provided for heating of the first zone works with
10 greater corrective action of manipulated value while the regulator of the second zone works with smaller corrective action of manipulated value. In other words, the temperature of the contents in the first zone may be changed more quickly and accurately than the temperature of the liquid in the second zone which has a larger area of heat transmission.

15 If a small volume is pipetted, it is quickly heated to the desired temperature. If a larger volume is pipetted, the pipette fluid is sucked into the second zone above the first zone, wherein the first zone acts as pretempering for the second zone. However, these volumes arrive in the second zone at temperatures that depend on pretempering that occurs in the first zone.
20 Generally, there will be a relatively low temperature differences of some degrees between the zones.

Due to the different structure of the two zones, temperature regulation of the second zone is more sluggish and is distinguished by small temperature fluctuations and by a larger area of heat transmission, in order to
25 be able to temper larger volumes in a short period of time.

It is preferable for the tube of the first zone to be relatively straight and for the tube of the second zone to be twisted or wound, for example, in either a spiral or in a meander configuration. This structure facilitates the smaller heat capacity of the first zone and the larger heat capacity of the second
30 zone.

It is also beneficial if the tube in the area of the second zone abuts a heat conductor, such as a metal block having an aperture for receiving the tube.

According to a preferred embodiment of the invention, the electrical
5 heater of the first zone is a PTC (positive temperature coefficient) heater wherein the heating wire has a varying resistance. Thus, if the tube in the first zone is only partly filled with the pipette fluid, the temperature of the PTC resistance wire in the unfilled part increases, which results in increased electrical resistance, so that the part that does not contact the fluid is heated
10 less. Due to this feature, a very even tempering may be obtained independent of the filling condition of the tube in the first zone.

A temperature sensor is provided at the entrance of the second zone so that the first temperature at the side of the entrance is measured.

Additionally, a similar temperature sensor also can be provided at the
15 egress of the second zone, which is beneficial for example for cleaning purposes, when the tube is filled in the reverse direction. For this reason, throughout the present application the zones are not referred to as lower and upper pipetting zones, but rather are referred to as a first and a second pipetting zone. Generally, the pipetting fluid is sucked through the tip to the
20 first, lower zone, and then arrives in the second, upper zone, if a sufficient volume is present.

In a preferred embodiment, in the second zone, the temperature sensor, and the heating element are identical.

The pipette tube according to the invention may also be used for
25 cooling. For this, the invention may include Peltier elements instead of the heating elements described above. Peltier elements, which are known, are used for both cooling or heating.

In a further preferred embodiment, the nominal temperature in the second zone is somewhat lower, preferably between 0.5 and 5° C lower than
30 in the first zone.

BRIEF DESCRIPTION OF THE DRAWING

The accompanying drawing, which is incorporated in and constitutes a part of this specification, illustrates one embodiment of the invention, and together with the description, seeks to explain the principles of the invention.

5 FIG. 1 is a schematic diagram of a preferred embodiment of the present invention.

DETAILED DESCRIPTION OF THE PREFERRED EMBODIMENT

A preferred embodiment of the invention will now be described with
10 further detail. FIG. 1 schematically illustrates a pipette tube according to the invention. The pipette tube includes a lower area 1 and a subsequent upper area 2. The lower area includes a pipette tip, and is characterized by a substantially linearly extending tube portion.

In the upper area 2, the tube may be spirally wound or may be wound
15 in other manners such as meander configurations. The tube of upper area 2 abuts a body 3, constructed of material with good heat conducting properties, so that the heat capacity is increased.

At least the largest length of the lower area 1 is heated by means of a first electrical heater 4 to which a regulator 5 is connected. Similarly, the
20 upper area 2 of the tube is also heated by means of a second electrical heater 6, to which a regulator 7 is connected.

Thus, the total length 1, 2 of the tube is heated by two separate heaters 4, 6, wherein two different regulators 5, 7 with different regulating characteristics are utilized. The regulator 5 works with a strong corrective
25 action of manipulated value and the regulator 7 has a smaller corrective action of manipulated value. In other words, the first heater has a superior ability than the second heater to quickly and accurately alter the temperature of the fluid in the tube to reach a predetermined temperature. There is a difference in the range of adjustments between the heaters due to differences
30 in the temperature changing magnitude of the regulators.

The first heater 4 heats the substantially linear portion of the tube up to, for example 15 mm before the tip. The second heater 6 includes an aluminum

cylinder (body 3) with a large heat capacity. The greatest part of the tube in respect to its length is wound around the cylinder.

The regulating circuit 5 of the lower heater 4 regulates, without delay, the temperature of the quantity of fluid in the lower area 1. This means that great fluctuations in the nominal size are regulated very quickly. If cold fluid is received, it is heated near the desired temperature when passing the lower part of the tube (area 1). If it is received in the upper part 2 of the tube which is tempered by the upper heater 6, the temperature is adapted quickly and without great fluctuations because of the smaller heat capacity of the fluid compared to the aluminum cylinder 3 and because of the small difference in temperature of the fluid to the temperature of the cylinder. Thus, a regulating circuit with very small corrective action of manipulated value is used.

The advantage of the described regulation is a very small fluctuation (change) of the temperature at the nominal value. The disadvantage of the longer regulation time for strong fluctuation of the nominal value thereby is not effective.

Tests have shown that with the pipette tube according to the invention, pipette fluids of, for example, 20 μm up to 600 μl can be pipetted wherein the different areas 1, 2 of the tube were utilized. In this case, the pipette fluid was heated within a few seconds to $37.3 \pm 0.3^\circ\text{C}$. The wall temperature of the tube should not exceed a temperature of 4°C in order to avoid damage to the pipette fluid.

FIG. 1 also shows a temperature sensor 8 at the inlet of the second zone. A further temperature sensor 9 may also be provided at the outlet of zone 2. The temperature sensors 8, 9 are part of the regulating circuit of regulator 7.

In a preferred embodiment of the invention, the lower area 1 has a volume of 100 μl and the upper area 2 has a volume of 500 μl . However, volumetric relationships other than a 1:5 ratio of the volume of the lower area to the upper area are possible.

Other embodiments of the invention will be apparent to those skilled in the art from consideration of the specification and practice of the invention

disclosed herein. It is intended that the specification and examples be considered as exemplary only, with a true scope and spirit of the invention being indicated by the following claims.

**THE EMBODIMENTS OF THE INVENTION IN WHICH AN EXCLUSIVE
PROPERTY OR PRIVILEGE IS CLAIMED ARE DEFINED AS FOLLOWS:**

1. A heatable pipette apparatus for heating and maintaining a liquid
5 conveyed in the pipette apparatus to a desired temperature, the pipette
apparatus comprising:
 - a pipette tube having a first zone and a second zone, the first zone
including a tip end;
 - 10 first heating means for heating the first zone;
 - second heating means for heating the second zone;
 - first and second regulating means for separately regulating the
temperature of the first heating means and the second heating means,
respectively, the first regulating means rapidly adjusting the
15 temperature of the liquid toward the desired temperature, and the
second regulating means finely adjusting the temperature to achieve
and maintain the desired temperature.
2. The apparatus as claimed in claim 1 wherein the tip end includes an
20 outlet of the pipette tube, and wherein a volume of liquid capable of being
held in the second zone is greater than a volume of liquid capable of being
held in the first zone.
3. The apparatus as claimed in claims 1 or 2 wherein the tip end includes
25 an outlet of the pipette tube, and wherein the first zone of the tube has a
substantially linear shape and the second zone of the tube has a twisted
shape.
4. The apparatus as claimed in any one of claims 1-3 wherein the second
30 heating means includes a heat conducting material abutting the second zone.

5. The apparatus as claimed in any one of claims 1-4 wherein the first heating means includes a PTC-heater.
6. The apparatus as claimed in any one of claims 1-5 wherein at least one
5 of said first and second regulating means includes a temperature sensor located between the first and second zones of the tube.
7. The apparatus as claimed in claim 6 wherein at least one of said first and second regulating means further includes a temperature sensor located
10 at an end of the second zone opposite the first zone.
8. The apparatus as claimed in any one of claims 1-7 wherein the first and second regulating means include respective first and second regulators, said first and second regulators being connected to the first and second
15 heating means, respectively.
9. The apparatus as claimed in any one of claims 1-8 wherein the first and second heating means include Peltier elements.
- 20 10. A heatable pipette apparatus for heating and maintaining a liquid conveyed in the pipette apparatus to a desired temperature, the pipette apparatus comprising:
- a pipette tube having a first zone and a second zone, the first zone including a tip end;
 - 25 first heating means for heating the first zone;
 - second heating means for heating the second zone;
 - first regulating means for controlling the first heating means; and
 - second regulating means for controlling the second heating means, the first and second regulating means being operable independently of
 - 30 each other.

FIGURE 1

