

included to effect this actuation along with a controlling algorithm for initiating sealing at the appropriate step in the analysis cycle.

[0176] The instrument also has an electrical connector of the type described in jointly owned It is used to make electrical connection to the sensor 128 in the housing 100. Where it is desirable to perform the detection step at a controlled temperature, e.g. 37° C., the connector also incorporates heating and thermistor elements, which contact the back side of the silicon chip that provides the substrate for the sensor. These elements are of the same type as described for the amplification chamber. The instrument has amperometric circuitry for controlling the potential of the sensor and measuring current. The instrument also has an embedded algorithm for controlling the entire analysis sequence performed by the instrument on the single-use device to make a nucleic acid determination and display a result on a display screen on the instrument. Where the electroactive species generated or consumed in proportion to the captured target is more appropriately detected by means of potentiometry or conductimetry, alternative circuitry well known in the art is incorporated into the instrument.

[0177] In an alternative embodiment, the single-use device is composed of two separate parts as shown in FIGS. 19 and 20. FIG. 19 illustrates a separate extraction device 470 and a combined amplification and detection device 471. The elements in a combined form have the same features as those shown for the integrated device in FIG. 6, with the exception of features related to transferring extracted material from one to the other. Element 470 comprises an entry port 413, conduit 411, wash fluid 417 and waste chambers 418, a separation region 421, a terminal portion of the conduit 601 and an egress port 502 which mates with ingress port 502. It also has mating features 520 and 521 which match one or more opening 500 in 471. Element 471 has an amplification chamber 410, conduit 409, chambers 408, 409 and sensors 419, 420, exit conduit 405 and sealing feature 406. FIG. 20 is similar to FIG. 19, with the difference that it comprises a combined extraction and amplification component 472 and a separated detection component 473. The mating features are appropriately located between the two.

[0178] FIG. 18 shows an additional embodiment where a filter region 421 is integrated into a device that performs extraction, amplification and detection. Other elements are as for FIG. 19. FIG. 17(a) shows an optical detection-based single-use cartridge where an optical sensor is integrated into the device that is interrogated by a reflectance method. Light is generated by element 401 and interacts with sensor 403 and is captured by detector 400. FIG. 17(b) shows an optical single-use cartridge where the sensing region is a cuvette feature 404, permitting detection with a light source 402 and detector 400 integrated into the instrument.

[0179] It has been found that where the sample is a buccal swab, the extraction component element, either magnetic or filter based, is unnecessary and the sample may be directly inserted into the amplification chamber. FIG. 28(a) and FIG. 28(b) show two views (top and bottom) of a buccal sample device for direct application of a buccal sample to a per chamber. This extraction and amplification device attaches to the detection cartridge, by means of the mating features described above (not shown).

[0180] The general dimensions of the housing 100 are about 6 cm in length, 3 cm in width and 0.3 cm in height. The conduits and other features are preferably rendered in a

device base 143 and a device cover 144 which are held together by an intervening double-sided adhesive tape 145, see FIG. 6. Where the base and cover are injection molded in plastic, typically ABS or polycarbonate, conduits and recesses to accommodate silicon chips, fluid containing pouched and the like are molded features. In this embodiment the adhesive tape acts as a sealing gasket to confine liquids to the desired conduits and chambers. Detailed discussion of the use of molded cover and base elements along with the use of adhesive tape gaskets is found in jointly owned U.S. Pat. No. 5,096,669 and pending US 20030170881 which are incorporated here by reference.

DETAILED DESCRIPTION OF DETECTION

[0181] The preferred method of detection in the single-use cartridge is electrochemical, however other sensing methods including fluorescence, luminescence, colorimetric, thermometric, fiber optics, optical wave guides, surface acoustic wave, evanescent wave, plasmon resonance and the like can be used.

[0182] The preferred sensor 128 comprises an amperometric electrode 300, which is operated with a counter-reference electrode 301 and is shown in FIG. 6. The amperometric electrode 300 comprises a 100 um diameter gold layer microfabricated onto a silicon chip 302. The silicon chip is treated in the first step of manufacture to produce an insulating layer of silicon dioxide on the surface, as is well known in the art. The electrode is connected by means of a conducting line 303 to a connector pad 304 which makes contact with the electrical connector of the instrument. The conducting line is typically coated with an insulating layer of polyimide 305. Directly over the electrode 300 or at an adjacent location 306 on the chip are adhered polymer particles 307 that have a ligand 308 complimentary to and capable of capturing the amplified target. The counter-reference electrode may be microfabricated on the same silicon chip or one place adjacently in the second conduit 125. It comprises a silver-silver chloride layer, of 200 um diameter attached by a contact line 309 to a contact pad 310 that makes contact with the instrument connector. Again the line 309 is preferably coated with an insulating layer of polyimide. A detailed description of amperometric sensor microfabrication is found in jointly owned U.S. Pat. No. 5,200,051 which is incorporated here by reference.

[0183] A conductivity sensor comprising two conductive bars 311 and 312 are present on chip 302, or an adjacent chip 350, connected to contact pads 313 and 314 by lines 315 and 316 respectively, see FIG. 6. The conductivity sensor can be used by the instrument to distinguish if liquid or air is in contact with the sensor and thus determine the position of a solution in the second conduit with respect to the sensor 300. This solution may be one containing the amplified target or the detection reagent. Optionally a conductivity sensor may be incorporated into or adjacent to both the extraction chamber and the amplification chamber to determine the position of a fluid. A detailed description of conductivity sensor microfabrication and use is found in jointly owned U.S. Pat. No. 5,447,440 and U.S. Pat. No. 6,750,053 which are incorporated here by reference.

[0184] In an alternative embodiment of the single-use device 100 a transparent glass window is substituted for the silicon chip 302 and the sensing region of the device forms a cuvette, FIG. 17. The amplified target capture reagent is immobilized on the glass and in this case the detection reagent