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(71) Applicant (for all designated States except US): MI-CRONICS, INC. [US/US]; 8463 154th Avenue Northeast, Redmond, Washington 98052 (US).

(72) Inventors; and

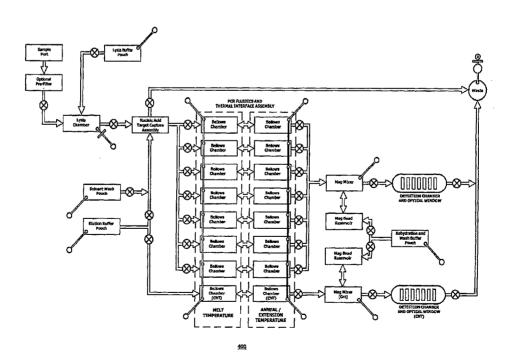
(75) Inventors/Applicants (for US only): BATTRELL, C. Frederick [US/US]; 18247 Northeast 28th Street, Redmond, Washington 98052 (US). GERDES, John [US/US]; 11 Club Lane, Columbine Valley, Colorado 80123 (US). WILLIFORD, John R. [US/US]; 18718 Northeast 55th Street, Sammamish, Washington 98074 (US). HOEKSTRA, Denise Maxine [US/US]; 16322 224th Street Southeast, Monroe, Washington 98272 (US). BREIDFORD, Wayne L. [US/US]; 5027 46th Avenue

South, Seattle, Washington 98118 (US). MORDUE, Stephen [US/US]; 254 4th Avenue, Apt. 3, Kirkland, WA 98033 (US). CLEMMENS, John [US/US]; 2554 174th Avenue Northeast, Redmond, Washington 98052 (US). NABAVI, Melud [US/US]; 2040 Waverly Place North #104, Seattle, Washington 98019 (US). KOKORIS, Mark [US/US]; 10535 Northeast 196th Street, Bothell, Washington 98011 (US).

- (74) Agents: WAGNER, Emily, W. et al.; Seed Intellectual Property Law Group PLLC, Suite 5400, 701 Fifth Avenue, Seattle, Washington 98104-7064 (US).
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(54) Title: INTEGRATED NUCLEIC ACID ASSAYS



(57) Abstract: Integrated microfluidic cartridges for nucleic acid extraction, amplification, and detection from clinical samples are disclosed. The devices are single-entry, sanitary, and disposable. The devices enable simplex or multiplex nucleic acid target detection, as for example: assay panels for multiple infectious agents, or assay panels for cancerous cell types. Methods for use of microfluidic cartridges in a fully automated, pneumatically controlled apparatus are also disclosed.

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ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, MT, NL, PL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

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A. CLASSIFICATION OF SUBJECT MATTER INV. C12Q1/68 B01L3/00							
According to International Patent Classification (IPC) or to both national classification and IPC  B. FIELDS SEARCHED							
	DEARCHED  Cumentation searched (classification system followed by classification)	tion symbols)					
C12Q B01L							
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 practical, search terms used)							
EPO-In	ternal						
C. DOCUMENTS CONSIDERED TO BE RELEVANT							
Category*	Citation of document, with indication, where appropriate, of the re	elevant passages	Relevant to claim No.				
Χ ,	US 5 587 128 A (WILDING PETER [U	IS] ET AL)	1,2				
	24 December 1996 (1996-12-24)	ing 35					
	column 4, line 10 - column 10, line 35 figure 12; example 2						
	column 13, lines 9-65						
	column 14, lines 8-62 column 18, line 61 - column 20, line 12 column 22, line 19 - column 23, line 27 column 25, lines 12-24						
	column 3, line 37 - column 4, line 6						
χ	WO 2005/106024 A (SIEMENS AG [DE	:];	1,2				
	GUMBRECHT WALTER [DE]; PAULICKA	PETER					
	[DE]; STANZEL M) 10 November 2005 (2005-11-10)						
	page 5, line 16 - page 6, line 25; figure						
	1						
Further documents are listed in the continuation of Box C. X See patent family annex.							
* Special c	ategories of cited documents:	"T" later document published after the inte					
A' document defining the general state of the art which is not cited to understand the principle or theory underlying the							
"E" earlier document but published on or after the international  "X" document of particular relevance; the claimed invention							
"L" document which may throw doubts on priority claim(s) or involve an inventive step when the document is taken alone							
citation or other special reason (as specified)  cannot be considered to involve an inventive step when the							
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"P" document published prior to the international filing date but the art. later than the priority date claimed "&" document member of the same patent family							
Date of the actual completion of the international search  Date of mailing of the international search report							
2	1 September 2007	18/01/2008					
Name and n	nailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2	Authorized officer					
NL – 2280 HV Rijswijk							
	Fax: (+31-70) 340-3016	Tiede, Ralph					

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## INTERNATIONAL SEARCH REPORT

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:
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 additional 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 an additional fees, this Authority did not invite payment of additional fees.
As only some of the required additional search fees were timely paid by the applicant, this international search reportcovers 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 (part), 2
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.

### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1(part), 2

A microfluidic cartridge comprising a fluidic subcircuit for extracting nucleic acids from a biosample, a fluidic subcircuit for synthesizing amplicons, a fluidic subcircuit with means for detecting amplicons and wherein said fluidic subcircuits are formed within a single integrated member, further comprising a sanitary waste receptacle

2. claims: 1 (part), 3,

A microfluidic cartridge comprising a fluidic subcircuit for extracting nucleic acids from a biosample, a fluidic subcircuit for synthesizing amplicons, a fluidic subcircuit with means for detecting amplicons and wherein said fluidic subcircuits are formed within a single integrated member, further comprising a multiplex detection chamber with a test chamber comprising antibodies bound to test pads

3. claims: 1(part), 4,26-37,50

A molecular detection complex and devices and methods comprising such a complex which comprise a two-tailed amplicon and wherein said complex is immobilized on a solid phase

4. claims: 1(part), 5-25, 38-49, 51

A microfluidic cartridge comprising a fluidic subcircuit for extracting nucleic acids from a biosample, a fluidic subcircuit for synthesizing amplicons, a fluidic subcircuit with means for detecting amplicons and wherein said fluidic subcircuits are formed within a single integrated member, further multiple channels and chambers for processing a sample and internal pneumatic control elements comprising diaphragm operated valves and bellows chambers

## INTERNATIONAL SEARCH REPORT

Information on patent family members

International application No PCT/US2007/006584

Patent document cited in search repo		Publication date		Patent family member(s)	Publication date
US 5587128	A	24-12-1996	US.	5955029 A	21-09-1999
WO 20051060	24 A	10-11-2005	CN DE EP US	1950520 A 102004021780 A1 1740722 A1 2007219366 A1	18-04-2007 24-11-2005 10-01-2007 20-09-2007