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di Studi Universitari e Perfezionamento



www.bioroboticsinstitute.eu

BIOSTEC 2014

7TH INTERNATIONAL JOINT CONFERENCE ON BIOMEDICAL ENGINEERING SYSTEMS AND TECHNOLOGIES

> Angers, France March 4, 2014

FRONTIERS OF SURGICAL ROBOTICS

Cesare Stefanini

The BioRobotics Institute Scuola Superiore Sant'Anna Pisa, Italy



The University and Research System in Pisa



PISA

SCUOLA NORMALE SUPERIORE

Scuola Normale **Superiore** Established 1810 150 professors 283 students





University of Pisa Established in 1343 2.100 professors 50.000+ students

UNIVERSITÀ DI PISA



Scuola Superiore Sant'Anna Established in 1987 100 professors 220 students



CNR National Research Council 15 Research Institutes About 1,500 researchers

NEST

National Enterprise for nanoScience and nanoTechnology

INFN National Institute of Nuclear Physics About 300 researchers



Tuscany: the Land of Robot(ic)s







This and more at: www.bioroboticsinstitute.eu





This and more at: www.bioroboticsinstitute.eu











The Robot

DustCart ATTENZIONE

attraversament





orog















Lamprey and Salamander Robots and the Central Nervous System





Biol Cybern (2013) 107:495-496 DOI 10.1007/s00422-013-0570-6

EDITORIAL

Biological Cybernetics

Foreword for the special issue on Lamprey and Salamander Robots and the Central Nervous System

Auke Jan Ijspeert · Sten Grillner · Paolo Dario









The intersegmental phase-lag - graded modification by action on a few rostral segments





Kozlov et al, PNAS, 106:20027-32, 2009



Brain circuitry





Transformée

Tectal Control of Locomotion, Steering, and Eye Movements in Lamprey Kazuya Saitoh, Ariane Ménard and Sten Grillner JNeurophysiol 97:3093-3108, 2007. First published Feb 15, 2007; doi:10.1152/jn.00639.2006



Results published on Bioinspiration and Biomimetics (vol. 7, p. 1-8, 2012)



Goal driven locomotion through vision

Colour tracking

Continue targeting

Random spot targeting



Results in press on <u>Biological Cybernetics</u>



Medical Robotics: an increasingly successful clinical and industrial field































Achievements of Robotics Surgery

- □ Game-changing applications
- Technically advanced and dependable systems
- Widely accepted and used in clinical practice by surgeons, by patients and by hospital administrations: 450.000+ surgical interventions worldwide in 2012
- Real IMPACT on health, and on economy (real products, real jobs)





The "Secrets" of the DaVinci Robot Success: Accuracy, Dexterity, Intuitiveness



- Outstanding mechanical design
- Excellent optics(2D and 3D vision)
- Smart and friendly interfaces

Successful daVinci applications in surgery





Open Hysterectomy Incision

da Vinci Hysterectomy Incisions





Open Prostatectomy Incision

da Vinci Prostatectomy Incisions

da Vinci® Hysterectomy Procedure Growth



Da Vinci Robot and Intuitive Surgical at-a-glance

South Korea 33

Japan 20 China 16

> aiwan 7 ingapore

Thailand 2

hiliopines

India 7



2.700+ systems installed in 30 Countries and in 1800+ hospitals

DaVinci Worldwide Installations

Bravil

Mexico 2 Chile 1

Argentina 3

Columbia



13

		Revenue		
	Year Ended December 31,			
2011	2010	2009	2008	29
(In	millions, except	per share amou	nts and headcou	int)
\$1,757.3	\$1,413.0	\$1,052.2	\$ 874.9	5

In 2012, 450.000 surgical procedures were performed with the da Vinci Surgical System. 1.500.000 procedures in total so far.

SOURCE: www.intuitivesurgical.com

The Evolution of Surgery



Why a Change of Paradigm is INEVITABLE

The operating room of the year 2030 will be a totally different environment than today

Professor Sir Alfred Cuschieri, MD

Scuola Superior Sant'Anna

THE BIOROBOTICS

Director of the Institute of Medical Science and Technology in Dundee and St Andrew's Universities. Pioneer of endoscopic surgery <u>MASS Screening and EARLY</u> <u>diagnosis</u> will have a major impact on the <u>type and</u> <u>invasiveness</u> of required surgical procedures

The combination of micro/nano/bio technologies, molecular biology, chemistry, physics, robotics/microrobotics, etc. will be key technologies enabling future high quality (accurate and repeatable), early and minimal invasive surgery

Prevention: the challenge of modern medicine



What is Endoluminal Surgery?

Endoluminal procedures consist of bringing a set of advanced therapeutic and surgical tools to the area of interest by navigating in the *lumina* of the human body, such as the gastrointestinal tract, the urinary apparatus, the circulatory system, etc., with a scarless or minimally invasive access



PillCam for GI tract endoscopy

Instrumentation for endoscopic surgery and NOTES (Natural Orifices Transgastric Endoscopic Surgery)



Clip for endoscopic surgery



The Neuroendoscope: an example of new endoluminal ultraminiature tool



NeuroEndoscopy of the spinal cord the Problem



There is a need for:

- Precise and early diagnosis of spinal cord lesions (300.000 paralyzed persons in Europe)
- Possibility to intervene directly on pathologies:
 - Injection of neurotransmitters and pumping of haematomas in traumatic lesions
 - Electro-coagulation of the afferences to the posterior horn in case of intractable pain
 - Cleaning of fibrous adherences in case of arachnoid proliferation

Current procedures are limited to the <u>epidural space</u>, far from the spinal cord and filled by semiliquid, **not transparent** fat.

An Endoscopic System is needed for the <u>navigation</u> inside the <u>subarachnoid space</u> (filled by the Cerebro Spinal Fluid), for diagnosis and intervention.



Neuroendoscopy of Spinal Cord The environment



- The mean is suitable for navigation (Cerebro-Spinal Fluid, water-like liquid)
- The workspace is extremely small (few millimeters)







Neuroendoscopy of Spinal Cord The environment



- The mean is very suiable for navigation (Cerebro-Spinal Fluid, water-like liquid)
- The workspace is extremely small
- Anatomical structures are very DELICATE (vessels, nerve roots)



AXONAL DAMAGE



For cord tissue, shear strains above 15% represent severe injury.









MINOSC – The Actuated Catheter

The concept:

• **Mechanical Steering** of the endoscope using <u>flexure</u> <u>joints</u> fabricated via Graded Material Technology and Injection Molding.





Servoassisted actuation

for fine and controlled advancement of the endoscope and steering. Servomotor actuating a STORZ micromanipulator.





MINOSC – The Fluidic System

Fluidic navigation system of

a microendoscope exploiting *microjets,* in order **not to touch** the tissue



Vision system:

0.5 mm OD endoscope, 6000 pixels + illumination







Automatic segmentation of lumen images



Nerve root

Pia Mater with blood vessel

Lumen



Pia Mater

nerve root with blood

Nerve root

Pia Mater with blood vessel





In vivo navigation







In-vivo validation

In vivo experiments in pigs in Ozzano (Bologna).

Successful endoscopy (see images below) of the whole spinal cord, from lumbar access up to cervical tract, with <u>direct</u> <u>nerve stimulation through</u> <u>endoluminal electrode</u>





Nerve root

Pia Mater with blood vessel



From **bio-inspiration** to **bio-application** (the EU FET BIOLOCH and the EMIL IMC Projects)



Problems in colonoscopy: pain difficult maneuverability...





Semi-autonomous inchworm-like locomotion







Retraction







WORM Painless Colonoscopy System

The E-






Professional

Partner

Careers

Int J Artif Organs. 2009 Oct 21;32(8):517-527. [Epub ahead of print]

Functional evaluation of the Endotics System, a new disposable self-propelled robotic colonoscope: in vitro tests and clinical trial.

Cosentino F, Tumino E, Rubis Passoni G, Morandi E, Capria A.

Gastroenterology and Digestive Endoscopy, San Giuseppe Hospital, Milan - Italy.

Abstract

Objective: Currently, the best method for CRC screening is colonoscopy, which ideally (where possible) is performed under partial or deep sedation. This study aims to evaluate the efficacy of the Endotics System, a new robotic device composed of a workstation and a disposable probe, in performing accurate and welltolerated colonoscopies. This new system could also be considered a precursor of other innovating vectors for atraumatic locomotion through natural orifices such as the bowel. The flexible probe adapts its shape to the complex contours of the colon, thereby exerting low strenuous forces during its movement. These novel characteristics allow for a painless and safe colonescency thus eliminating all major associated risks such as infection, endispumonary complications and colon perforation. Methods: An experimental study was devised o investigate stress pattern differences between traditional and robotic colonoscopy, in which 40 enrolled patients underwent both robotic and standard colonoscopy within the same day. Results: The stress pattern related to robotic colonoscopy was 90% lower than that of standard colonoscopy. Additionally the robotic colonoscop, demonstrated a higher diagnostic accuracy since, due to the lower incomation rate, it was able to visualize small polyps and angiodysplasias not seen during the standard colonoscop, All patients ared the robotic colonoscopy as virtually painless compared to the standard colonoscopy, ranking pain and discomfort as 0.9 and 1.1 respectively, on a scale of 0 to 10, versus 6.9 and 6.8 respectively the standard device. Conclusions: The new Endotics System demonstrates efficacy in the diagnosis of colonic pathologic using a procedure nearly completely devoid of pain. Therefore, this eveter wan also be looked upon as the first step toward developing and implementing colonoscopy with atraumatic locomotion through the bowel while maintaining a high level of diagnostic accuracy.



From "<u>wired</u>"painless colonoscopy to "<u>wireless</u>" GI endoscopy





Wireless endoscopic capsules with <u>active</u> <i>locomotion system for the entire GI tract



A. Moglia, A. Menciassi, P. Dario, A. Cuschieri, "Clinical update: endoscopy for small-bowel tumours", THE LANCET, Vol 370 July 14, 2007, pp. 114-116

M. Quirini, S. Scapellato, A. Menciassi, P. Dario, F. Rieber, C.-N. Ho, S. Schostek, M.O. Schurr, "Feasibility proof of a legged locomotion capsule for the GI tract", GASTROINTESTINAL ENDOSCOPY Vol. 67, No. 7, 2008

ACTIVE capsules developed at The BioRobotics Institute





Wireless Capsule for PAINLESS GASTROSCOPY

Ingestion of liquid in context with the examination allows to obtain organ distension, thus making possible a low power 3D locomotion in the stomach



Wireless Capsule for PAINLESS GASTROSCOPY

D- DB	Swimming Capsule Control		
Stop motori	Video	State ON Error Msy 10 • • • • • • • • • • • • • • • • • • •	
STOP POWER OF OF STATES	N D O O D O O O O O O O O O O O O O O O	PWH 4 PWH 3 0 0 0 PWH 1 PWH 2 0 0	
Motor 2 Hotor 1 50 50 Motor 3 Hotor 4 50 50	Trané Rate Drage Aurice 20,455 230 352k240 (2: 32*0r RGB mage 130, 155,225* (0,0)	100 80 00 40 20	
CET BATTERY VOLTACE		a²∎	

How to overcome the energy problem in active capsular endoscopy: magnetic assisted locomotion



Endoluminal magnetic locomotion can be extremely precise when the external magnet (s) is/are moved by means of high precision robot(s)





Custom image sensor









Scuola Superiore Sant'Anna









Resolution		QVGA
Active area		320×240
Optical format	inch	1/11
Pixel Pitch	μm^2	4.4×4.4
Shutter type		rolling
Die dimension	mm ²	2.5×3.0

Sensitivity	lux	$0.1@555nm,27^{\circ}$ C, 30msec.
	W/m^2	$1.7 \times 10^{-4} @ 27^{\circ}$ C, 30msec.
Responsivity	V/lux*sec.	$0.52@555$ nm, 27° C
	$V/W/m^{2*}sec.$	360@@27°C
Dynamic range	dB	50
SNR	dB	46(max)
Pixel Noise	%	0.70
Fixed Pattern	%	0.86
Noise		

A Novel Concept: Magnetically-Controllable Insulin-Filled Capsules as Carriers for Diabetes Treatment

- □ 347 million people worldwide suffer from diabetes
- □ diabetes deaths are expected to double between 2005 and 2030
- □ worldwide diabetes market: from \$ 14.9 billion in 2009 to \$ 55 billion in 2019
- Iifestyle strongly affected by traditional therapy



Multiple daily insulin injections

- The patient is slave of his/her pathology
- Long-term complications
- Non-physiologic insulin profile

Implanted artificial pancreas

- Limited lifetime
- Need of complicated periodical refilling procedures (surgical operations needed every 3 months)
- Low insulin stability in the implanted device



Wearable artificial pancreas

- Delays in insulin adsorption
- Common daily activities strongly affected



Holy Grail of diabetes treatment Long-term totally implantable AP refillable through insulin pills



Novel implantable artificial pancreas *



From **Single** Capsules to a **Molteplicity** of Capsules: **Modular** and **Reconfigurable** Surgical Instruments











THE BIOROBOTICS

Scarless Robotic Surgery

Laparoscopic Procedure















The EU ARAKNES Project: www.araknes.org

The ARAKNES System Architecture



SPRINT: Single-Port lapaRoscopy bImaNual robot





The ARAKNES Project has received funding from the European Community's Seventh Framework Programme (FP7/2007-2013) under grant agreement num. 224565.

ARAKNES Project MNBS 2012 – Athens – 3th May 2012





The SPRINT robot



Single Port External Diameter: 30 mm

External Diameter of the robot arms: less than 20 mm



The ARAKNES Project has received funding from the European Community's Seventh Framework Programme (FP7/2007-2013) under grant agreement num. 224565.





R.A.K.N.E.S. In-Vivo Tests: Results

Small bowel enteroenterostomy



Ligation of a mesenteric vessel bundle



Results of In-Vivo Tests presented at **SAGES** Annual Meeting:

A New Robotic System for Single-Incision Laparoscopic Surgery: Preliminary Experience.

G. Basili, G. Pietroni, A. Menciassi, D. Pietrasanta, M. Niccolini, O. Goletti







Biological and bioinspired manipulators





The key aspect: to manipulate objects while controlling the stiffness of selected body parts and being inherently compliant when interacting with objects

Muscular Hydrostat of tentacles, trunks, and tongues: **modifiable skeleton** which allows the transformation of force into motion







OCTOPUS arm 18 SMA modules 12 tendons No rigid parts



The STIFF-FLOP modular arm

Overall



Architecture consisting of connected modules



Overall performance

The concept design of a new modular manipulator for MIS has been presented

One single module was fabricated and characterized, which can:

- Bend in any direction up to 120°
- Elongate up to 86.3%
- Stiffen locally up to 36%
- Produce force 47N of
- 47N of force at tip
- Squeeze for fitting into small diameter holes up to -40%

...towards the modular manipulator with 3+ modules

Pneumatic tubes



Stiffening vacuum line

WWW.STIFF-FLOP.EU









An endoluminal robot could also be seen as a carrier of micro/nano therapy agents (i.e. a SHUTTLE)





Some examples of nanotechnologies contributing to endoluminal and cell surgery

Exploitation of chemical and physical properties of nanomaterials in endoluminal and cell surgery

Carbon nanotubes (CNTs): blending physical and chemical properties for electroporation, localized hyperthermia, magnetic guided drug delivery, DNA transfection

Boron nitride nanotubes (BNNTs): from nanotubes to nanotransducers! Enhanced physical properties with the same chemical properties of CNTs

Magnetic nanofilms: merging the magnetic control with the therapeutic abilities of nanofilms in endoluminal surgery

Nano-machines for cell therapy: the NINIVE project (www.niniveproject.org, 6FP NMP 033378)

OBJECTIVE and NOVEL CONVERGING APPROACH:

combining physical and chemical properties of CNTs holds great promise for the development of a new class of CNT-based drugs and therapies extremely controlled (i.e. much more controlled that methodologies based on diffusion, phagocytosis, endocytosis)

Electrical properties (conductive and semi-conductive CNTs); optical properties; strong anisotropy; etc. **Covalent bonding of CNT surface; non-covalent absorption on the CNT surface; etc.**



Cell manipulation with magnetic carbon nanotubes

Once the CNTs (naturally magnetic thanks to residuals) are attached or internalized, cells can be concentrated in a desired compartment for subsequent localized therapy.

Tumor Tumor Magnet Metastatic cells bound with CNTs

Human Neuroblastoma cells (SH-SY5Y) displacement after 3 days in culture with MWNTs-modified medium. Control sample not showed (with Nikon TE2000U inverted optical microscope, magnifications 20x).

Cell

CNT can be functionalized to bind target cells (such as metastatic cells) or to be

become magnetotactic and can be drag and collected by a permanent magnet.

internalized by the cells; in this sense cells

V. Pensabene, O. Vittorio, S. Raffa, A. Menciassi, P. Dario, "Neuroblastoma cells displacement by magnetic carbon nanotubes", IEEE Trans. On NanoBioScience, 2008.



СИТ



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Perspectives related to PZT properties

The piezoelectric properties of BNNTs make them attractive candidates as **bionanotransducers**. If stimulated with non-invasive ultrasounds, they should be able to generate electrical field. We are carrying out experiments on cells sensible to electrical field (neurons, osteoblasts, muscle cells, etc.) and preliminary results on primary human osteoblasts show a significant increment of osteocalcine and calcium content after incubation with BNNTs and stimulation with US.





Polymer ultra-thin nanosheet for endoluminal surgery

In **NOTES** (Natural Orifice Transluminal Endoscopic Surgery), access to the target organs is obtained through holes made in stomach/female reproductive/lung wall.





Conventional technique to close a hole using endoluminal devices



Non invasive, flexible, efficient methods for hole closing are deeply investigated because current techniques show many limitations





Thanks to their flexibility, these nanofilms can be proposed as <u>nanoplasters</u> for closing incisions and wounding in endoluminal surgery procedures.
They can be stored in small channels of endoscope and can be delivered without loosing their flexibility.



Adding magnetic properties to the film, nanosheets can be precisely positioned in situ with catheters or robotic modules inside the stomach or other orifices.



Homogeneous Magnetic Nanosheets



Manipulation test

A Neodymium Iron Boron permanent magnet (Br= 350 mT) is used to move the film in saline solution and finally the film is controlled and attached on the tissue.

Videomagnetic manipulation

Attachment of nanofilms on stomach tissue



PLA 10 mg/ml, no particles





10 mg/ml NP (200 nm) in PLA 10 mg/ml

10 mg/ml NP (40 nm) in PLA 10 mg/ml

After the removal of the liquid, the film adhere on the surface and, thanks to the nanometric thickness, it precisely fits the morphology of the tissue.

Magnetic nanofilms as nanoplaster for endoluminal surgery

10 mg/ml NP (200 nm) in PLA 20 mg/ml



Film covering an incision on the mucosal wall

gastric wall

magnetic nanofilm



10 mg/ml NP (40 nm) in PLA 10 mg/ml



The film follows the folds of the mucosal wall, completely covering it.



Concluding remarks

Robots have a place in the modern operating room, because of their established ability – in a growing number of different fields - to exploit the increasing power of planning, imaging, diagnostics and teleoperation techniques to improve surgical outcome

The new frontier of medical robotics: miniaturization and integration of different functions

Enabling approach: the convergence of many disciplines, basically life science, physics, chemistry, neuroscience, nanotechnology, etc.





Acknowledgments

Many colleagues, many PhD students, many funding agencies (mostly the *European Commission*)



The BioRobotics Institute www.bioroboticsinstitute.eu

