PERSPECTIVES

The eminent need for an academic program in universities to teach nanomedicine

Juan Manuel Vélez Juan Jesus Vélez

Intracellular Signaling Lab, Sección de Estudios de Posgrado e Investigación, Escuela Superior de Medicina, Instituto Politécnico Nacional, México Abstract: Nanomedicine is on the cutting edge of technology applied to medical and biological sciences. Nanodevices, nanomaterials, nanoinstruments, nanotechnologies, and nanotechniques (laboratory methods and procedures) are important for the modern practice of medicine and essential for research that could stimulate the discovery of new medical advances. Accordingly, there is an eminent need for implementing an academic program in universities to teach this indispensable and pragmatic discipline, especially in the departments of graduate studies and research in the areas of pharmacology, genetic engineering, proteomics, and molecular and cellular biology.

Keywords: nanodevices, nanotechnologies, nanotechniques, nanobiosensors, quantum dots

Introduction

To be in step with the future, it is necessary today for academic programs to exist that teach topics related to nanomedicine, an important and relatively recent technological advance in medicine. Such programs should be included in university departments for graduate studies and research, especially in the areas of pharmacology, genetic engineering, proteomics, and molecular and cellular biology. Later, as nanotechnology continues to develop, other areas will certainly be affected.

Research in nanomedicine began with discoveries of the novel physical and chemical properties of various carbon-based materials that can only be detected in nanometer-sized structures. Nanomedicine is defined as the monitoring, repair, construction, and control of human biological systems at the cellular, molecular, and atomic levels using engineered nanodevices and nanostructures. Another definition is that provided by the National Institute of Health in its roadmap for medical research in nanomedicine, which states that nanomedicine is "an offshoot of nanotechnology, [which] refers to highly specific medical interventions at the molecular scale for curing disease or repairing damaged tissues, such as bone, muscle, or nerve".¹ Nanomedicine contains five main disciplines: analytical tools, nanoimaging, nanomaterials (medications, raw materials, and devices), novel therapeutic and drug delivery systems, as well as regulatory and toxicological issues related to clinical practice.¹

Basic nanostructured materials, engineered enzymes, and the many products of biotechnology will most probably become essential elements in some areas of medical, chemical, and biological applications. However, the full promise of nanomedicine is yet to be defined, as this will to a large extent depend on the development of new technology, such as precisely controlled or programmable medical nanomachines

Correspondence: Juan Manuel Vélez Laboratorio de Señalización Intracelular, Sección de Estudios de Posgrado e Investigación, Escuela Superior de Medicina, Instituto Politécnico Nacional, México Tel +1 55 57296000 ext 62824 Email jvelez@ipn.mx

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and nanorobots to address medical problems. Moreover, the application of nanotechniques at the cellular, molecular, and atomic levels should certainly help in the understanding of the function of organisms, as well as in the development of better techniques for imaging, diagnosing, and treating disorders at the level of molecular and cellular biology.

Microscopic machines were first hypothesized by the Nobel Prize winning physicist Richard P Feynman, now considered the father of nanoscience, in 1959. His article "There is Plenty of Room at the Bottom" is considered a classic in the establishment of the important possibilities for this field.² The main idea of this article was the ability to reduce the size of various apparatuses and at the same time increase their effectiveness, based on the manufacture of products by reordering atoms and molecules. Of particular importance in his proposal was the reference to nanorobots, to be used in the manufacture of nanodevices, nanomaterials, and nanomachines. He also mentioned the possibility of creating a 100-fold improvement in the electron microscope.

Eric Drexler, considered the father of nanotechnology, wrote the popular books, *Engines of Creation* (1986) and *Unbounding the future* (1991), in which he announced the promise and warned of the dangers of molecular and atomic manipulation. In his more recent work, *Nanosystems: molecular machinery, manufacturing and, computation* (1992), he predicted that nanotechnology will make the conventional means of manufacture obsolete.³ He proposed the concept of machines with the capability of ordering and reordering atoms in order to create precise molecular structures, thus resulting in the desired product. Such machines could even have the ability of self-reproduction.

Once nanomachines are available, programmable and controllable microscale robots comprised of nanoscale parts manufactured to nanometer precision will open new possibilities related to curative and reconstructive procedures in the human body at the cellular, molecular, and atomic levels (perhaps at less than 100 nm). In keeping with the adage that "the best treatments are those that intervene the least", the possibility of nanotechnology to greatly increase knowledge of cellular and atomic events could lead to improved treatments. Additionally, nanomedicine is on the cutting edge of medical practice and research, and for this reason it is an important focus of capital investment. Consequently, it is urgent that academic programs be implemented in order to teach this discipline in universities that have a medical and biological focus of study and research.

An academic discipline of nanomedicine at the university level would allow students, teachers, technicians, and researchers to become familiar with: (1) nanodevices, nanostructures, nanomaterials, nanostuffs, and precise and sophisticated nanoinstruments, such as those that could be used for molecular imaging and bedside biomarker testing and monitoring (Appendix 1); (2) the main categories of precise nanotechnologies in real time (Appendix 2); and (3) techniques (laboratory methods and procedures) needed for furthering research (Appendix 3), including detailed procedures of nanobiosensors for medical uses (Appendix 4). All of these elements of nanomedicine could be applied to cancer, cardiovascular diseases, toxicity, ethics, and societal and environmental impact.⁴

Perhaps one of the most important potentials for nanomedical research is the gathering of extensive information about the chemical and physical properties of nanoscale biological structures. As the catalog of the interactions between individual molecules, cells, tissues, and organs develops, researchers will gain a greater understanding of the intricate operations of molecular structures, processes, and networks inside living cells based on the use of new probing tools. Consequently, scientists will be able to build new devices for a wide range of biomedical applications, such as detecting infectious agents or metabolic imbalances with tiny sensors, generating miniature devices that search for and destroy infectious agents, and replacing "broken" machinery inside cells with new nanoscale structures.⁴

Current applications

The following medical advances are among the existing uses of nanomedicine.

- Nanomaterials: nanotechnology used for subcutaneous treatments.⁵
- **Nanoprobes in vivo:** targeted nanoprobes reveal early time point kinetics in vivo by time-resolved magnetic resonance imaging (MRI).⁶
- Methods for the preparation of small interfering ribonucleic acids (siRNAs): a direct, efficient method for the preparation of siRNAs containing ribo-like north pseudo-sugars,⁷ and drug delivery of siRNA therapeutics.⁸
- **Physical assessment of toxicology at nanoscale:** nanodose-metrics and toxicity factor.⁹
- Nanomaterials for regenerative medicine: use of nanomaterials with applications for regenerative medicine.¹⁰
- Vascular toxicity: studies of vascular endothelial toxicity.¹¹
- **Nanocatalizers:** molecular-level biological transporters capable of converting biomass into biofuel.

• **Nanoparticle devices:** molecular-level biological transporters that release medication in a precise and controlled manner throughout the organism, such as in target cells, for crossing biological barriers and for providing greater stability in the face of pathogenic deterioration, as well as use of nanoparticles for the treatment of cancer.¹²

Future directions and challenges

New opportunities will most certainly appear in relation to fighting disease as nanotechnology develops. Doctors may be able to search out and destroy the very first cancer cells before they lead to a destructive tumor in the body. Perhaps a broken part of a cell could be removed and replaced with a miniature biological machine. Pumps the size of molecules could possibly be implanted to deliver life-saving medicines precisely when and where they are needed. These scenarios are the long-term goals of the "nanomedicine roadmap".⁴ With the inclusion of the academic discipline of nanomedicine in their programs, universities will be able to provide the personnel capable of accelerating the development of this field. Future possibilities include the following opportunities:

- **Nanoneedles:** the nucleus of live cells could be penetrated by nanoneedles used in conjunction with microscopy with the aim of introducing nucleic acids, proteins or other compounds inside human cells, or even doing surgery; modeling the structure-property relationships of nanoneedles.¹³
- Environmental nanocatalizers: biological transporters capable of greatly improving catalytic converters currently used in automobiles, thus drastically reducing the pollution emitted.
- **Nanobiosensors:** sensors capable of detecting the presence of anthrax and other biological weapons; nanosensors for probing live cells.¹⁴
- **Dendrimers:** artificial molecules designed to replace defective endogenous elements in the organism, induce the production or secretion of various molecules related to the immune response in case of being genetically deficient, or act as biological markers (for cancer therapy, for the detection of adherence to medication regimens, or for the evaluation of experimental results).¹⁵
- Nanodepots of insulin: microscopic deposits injected into the bloodstream that could act as an artificial pancreas, releasing insulin to counteract the effects of diabetes; similar technologies could also be employed to release dopamine into neurons as therapies for Parkinson's or Alzheimer's disease.
- **Titanium dioxide nanotubes:** microscopic tubes that could clean polluted water.

- **Carbon nanotubes:** microscopic tubes aimed at replacing defective or absent genes; use of carbon nanotubes in vivo to characterize action mechanisms at the cellular level;¹⁶ carbon nanotubes could reactivate synaptic connections and regenerate neurons damaged by an accident.
- **Biological nanosensors:** a set of ultramicroscopic and fine silicon threads.
- Molecular nanopyramids: these probes could possibly be released into the human body, with the capability of detecting and destroying the first cancer cells in the brain, heart, kidney, or lungs, without damaging neighboring tissue; they might also be able to detect viral or bacterial infections in an early stage before the immune system has begun to respond.
- Assembler nanomachines: machines based on the rearrangement of atoms and molecules to manufacture molecular components.
- Smart materials: these probe-like materials could potentially detect subtle changes in body chemistry.
- **Nanoparticles:** microscopic tumor-destroying genes may possibly be transported to certain sites, directed with great precision to destroy the genetic information of tumors, with the aim of inactivating the target cells and impeding their reproduction; other genes may be transported to detect tiny alterations in proteins, as well as to scan tissue to ensure that the nanodevices are neatly congregated around the target.
- **Nanorobots:** these devices could be introduced into the human body to repair damage and combat infection and disease by acting on DNA without being detected and destroyed by the host immune system.^{17–22}
- **Nanobiochips:** this advanced technology could work with MRI to supply an abundant amount of information about the human body.²³
- **Nanocomputers:** computers could be developed made of material stronger than steel but with only 10% of its weight. These could possibly guide nanomachines for the purpose of examining, taking apart, and rebuilding damaged cells, tissues, and organs through the reconstruction of their molecular structures; nanoinformatics could be applied to nanomedicine;²⁴ new computers could search for a methodology that reproduces human organs in vitro.

Conclusion

The academic discipline of nanomedicine could be included in university curriculums for graduate students and researchers today in the fields of pharmacology, genetic engineering, proteomics, and molecular and cellular biology. Such programs would train personnel in order to accelerate advances of techniques, technology, and applications of nanomedicine. As such uses increase and broaden, other new programs will also become important elements of university curriculums.

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Disclosure

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Appendices

Appendix I Nanomateria	ls, nanodevices	, nanoinstruments,	and nanostuffs	useful in nanomedicine
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Nanostructures	Nanoparticles	Nanofibers	Nanoplatforms
Nanodisks	Nanoscaffolds	Nanolabels	Nanodiamonds
Nanoarrays	Nanocomposites	Nanopackages	Nanobubbles
Nanobeads	Nanofluidics	Nanobiosensors	Nanogadgets
Nanofilaments (silicone)	Nanotargets	Nanofilms	Nanoscale
Nanotubes (carbon)	Nanorods	Nanoshapes	Nanocoatings
Nanosuspensions	Nanoemulsions	Nanocarriers	Nanospheres
Nanosystems	Nanogels	Nanocontainers	Nanoproducts
Nanodrugs	Nanovectors	Nanohybrids	Nanoreactives
Nanoscopic	Nanoclusters	Nanobullets	Nanoknittings
Nanoswitches	Nanoblobs	Nanohazards	Nanobodies
Nanosurfaces	Nanopores	Nanocatalysts	Nanotools
Nanochips	Nanorobots	Nanocrystals	Nanoneedles
Nanowires (silicone)	Nanoshells	Nanoencapsulation	Nanoprobes
Nanolayers	Nanomotors	Nanomachines	Nanocages
Nanoimpellers	Nanomatrix	Pharmacytes	Dendrimers
Nanoscopic weapons	Qubit	SPION	Quantum dots

Abbreviation: SPION, superparamagnetic iron oxide nanoparticles.

Appendix 2 Technologies

		Appendix 5 (continued)		
MIDD: molecular imaging drug delivery	Artificial engineering	Tissue specimen preparation	PCR-based detection	
Nanoparticle drug delivery systems	Nanoscopic medicine	QD fluorescent nanoprobes	QD-antibody conjugation	
Tissue engineering regenerative medicine	Nanodimensional three	Image guided drug delivery	Lasser nanoscissor	
Cell therapy regenerative medicine	Cell surface engineering	Fluorescence resonance energy transfer	Cryopreservation	
UMI: ultrasound molecular imaging	Nanodiagnostics	Cryogenic transmission electron	Electrontomography	
Wireless endoscopy and biotelemetry	Regenerative medicine	microscopy		
AFM: atomic force microscopy	Nanofluidic technology	SLN: solid lipid nanoparticles	Fluorescence nanoscopy	
Nanoparticles for molecular diagnostics	Stem cell tissue engineering	Magnetic resonance propulsion	Colorimetric method	
Stem cell repair and therapy	Nanocybernetics	Electrostatic droplet generation	Nanoencapsulation	
Stem cell therapy for cancer	Laser nanoscissor	Molecular imprinted polymer particles	Radioimmunotherapy	
Nanoparticles for tracking stem cells	SPR: surface plasmon	Affinity nanoliquid chromatography	UV-bioimprint technique	
	resonance	Transmission electron microscopy	Magneto electroporation	
Nanoscale single-cell or molecule	Molecular diagnostics	Contrast imaging technique	BioMEMS	
identification		Nanomechanical holography	Nanophotothermolisis	
Nanotechnology-based cytogenetics	Endothelial cell repair	Plasmonics and SERS nanoprobes	Biomarker quantification	
Cancer molecular imaging	Nanoscale probes	Isolation and analysis of individual	MRI scanner	
Nanotechnology-based biochips and	RAMiTS technology	biomolecules		
microarrays		Nanolaser scanning confocal	Biobarcode assays	
Virtual reality stimulation	Nanoproteomics	spectroscopy	1	
Phototermical therapy	Organic electronic	Gold nanoparticles for diagnostics	Mass spectrometry	
Biodegradable polymeric nanoparticles as o	Irug delivery devices	A spectroscopic assay based on SERS	Standard	
Modular robotics, multi-agent or swarm robotics, and utility fog			photolithographic	

Appendix 3 Techniques and methods

QDs broad absorption spectra	Miniaturization top-down	
FISH: florescent in situ hybridization	Etching technique	
Atomic force microscopy and	The photostability	
quantum dot		
Scanning near-field optical microscopy	Fluorescence microscopy	
Functional polymer-coated nanoparticles	MRI scanner	
Lumininescent fluorescence microscopy	Confocal microscopy	
Immunicon's cell tracks technology	ELISA-based method	
	(Continued)	

Appendix 3 (Continued)

issue specimen preparation PCR-based detection			
QD fluorescent nanoprobes	QD-antibody conjugation		
Image guided drug delivery	Lasser nanoscissor		
Fluorescence resonance energy transfer	Cryopreservation		
Cryogenic transmission electron	Electrontomography		
microscopy			
SLN: solid lipid nanoparticles	Fluorescence nanoscopy		
Magnetic resonance propulsion	Colorimetric method		
Electrostatic droplet generation	Nanoencapsulation		
Molecular imprinted polymer particles	Radioimmunotherapy		
Affinity nanoliquid chromatography	UV-bioimprint technique		
Transmission electron microscopy	Magneto electroporation		
Contrast imaging technique	BioMEMS		
Nanomechanical holography	Nanophotothermolisis		
Plasmonics and SERS nanoprobes	Biomarker quantification		
Isolation and analysis of individual	MRI scanner		
biomolecules			
Nanolaser scanning confocal	Biobarcode assays		
spectroscopy			
Gold nanoparticles for diagnostics	Mass spectrometry		
A spectroscopic assay based on SERS	Standard		
	photolithographic		
Bioconjugated nanoparticle based bioassay			
Multicolor oligonucleotide-functionalized QDs			
SPION: super paramagnetic iron oxide nanoparticles			
Nano dissection and nanoextraction of chromosomal DNA			
Narrow emission spectra of nonorganic QD fluorophores			

Abbreviations: ELISA, enzyme-linked immunosorbent assay; MRI, magnetic resonance imaging; PCR, polymerase chain reaction; QD, quantum dot; SERS, surface-enhanced Raman scattering.

QDs for molecular diagnostics for cancer diagnosis and for viral diagnosis

Application of nanoparticles for discovery of biomarkers

Appendix 4 Nanobiosensors

Nanowire biosensors	lon channel switch biosensor
	technology
Cantilevers as biosensors	Electrochemical nanobiosensors
Carbon nanotube biosensors	FRET-based DNA nanosensor
Electronic nanobiosensors	Quartz nanobalance DNA sensor
Viral nanosensor	Nanosensors based on surface
	plasmon resonance
Optical biosensors using laser,	PEBBLE
nanoshell, SPR, SERS, mRNA	

Abbreviations: FRET, fluorescence resonance energy transfer; PEBBLE, probes encapsulated by biologically localized embedding; SERS, surface enhanced Raman scattering; SPR, surface plasmon resonance.

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