

**Next Generation Sequencing –  
The Role of New Sequence Technologies in Shaping the  
Future of Veterinary Science**

**Hosted by the RCVS Charitable Trust**





# The Future of sequencing

Neil Hall



# Sequencing...the new dot-com?

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### Roche raises offer for Illumina to over \$6bn

Roche has raised its offer for the gene sequencing firm Illumina.

The Swiss firm has raised its offer from \$44.50 per share to \$51, valuing the deal at more than \$6bn (£3.8bn).

Roche hopes the new bid will win over Illumina shareholders who gather for the annual general meeting next month.

The purchase of Illumina would help Roche build its personalised healthcare business, as Illumina's gene sequencing technology can identify which treatments suit different patients.

Illumina's management rejected Roche's initial offer made in January, so the Swiss pharmaceuticals giant has now taken its offer directly to shareholders.

"Based on our discussions with Illumina shareholders we have seen interest to accelerate the takeover process," said Roche chief executive Severin Schwan.



Roche is the world's biggest maker of cancer drugs

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[Glaxo invests £500m in UK sites](#)

- Sequencing is attracting a lot of investment even in an economic slump
- Driven by human medicine
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**The Best of 2011**

Even amid the chaos and uncertainty of the year, there were some bright and heroic moments. Here, we present seventy-seven things to celebrate.

By Andrew Chaikivsky

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15 of 77

Baffled by six-year-old Nicholas Volker's deadly intestinal disease, which forced him to eat through a feeding tube, doctors at Children's Hospital of Wisconsin in Milwaukee sequence the boy's genome in an effort to diagnose and cure him. **A medical first.** Identifying a never-before-seen mutation in his DNA, they surgically rebuild his immune system through a cord-blood transplant. After a 121-day hospital stay, Nicholas Volker returns home to enjoy his

# With Death, Christopher Hitchens And Steve Jobs Showed Us The Limits Of DNA Sequencing

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Christopher Hitchens, quite famously, did not believe in miracles. His death is a reminder that we shouldn't, either – even when they're the scientific kind.

Hitchens, like [Steve Jobs](#), was among the first patients to benefit from a very new technology: the use of DNA sequencing to pick cancer drugs that might have a better chance of slowing a tumor's growth.

Cells become cancerous because of mutations in their DNA that make them stop behaving as discrete parts of the body and instead cause them to multiply like crazy and run amok. Once a cell is cancer, its genes get twisted and re-arranged even more. The idea is that by



Image by Getty Images Europe via @daylife



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## The cancer revolution

When Sarah Allen, a mother of four, discovered she had a virulent form of breast cancer, she feared for her future. Two years on, she is the symbol of a biotechnological success that promises to change for ever our treatment of disease

Jo Revill  
 The Observer, Sunday 9 March 2003  
 Article history

Sarah Allen doesn't realise it yet, but she is at the forefront of a cancer revolution. The experience of this Midlands housewife gives us a glimpse into the future of medicine.

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#### DNA sequencing matches cancer patients to clinical trials

30 November 2011

A scanning machines could help identify cancer individual genetic make-up to match them to appropriate according to scientists in the US.

University of Michigan Comprehensive Cancer Michigan Centre for Translational Pathology (MCTP) ran exploring how to quickly and systematically sequence rial from patients with advanced or treatment-resistant

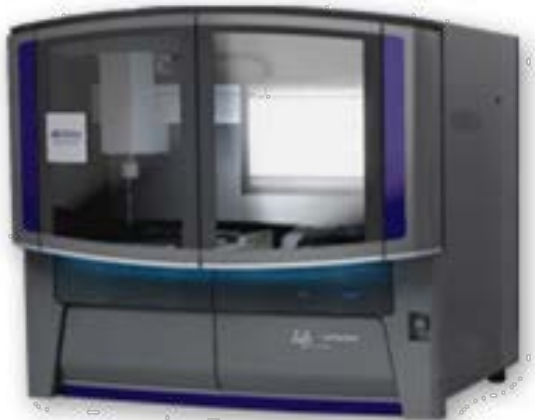
g DNA to spot which genes are faulty in a patient's tists can identify current clinical trials that include treatments targeted to these



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"We need more intelligent ways to match patients to clinical trials that are likely to be suitable for their particular cancer"  
 - Kate Law, Cancer Research UK

# Sequencing will get smaller



Personalised medicine needs personalised sequencers.

# Sequencing is moving out of centres

- MiSeq and Ion Torrent sequencing machines are appearing in non-specialist labs.
  - Simple(er) to use
  - Built in analysis tools
  - >1Gb throughput
  - cheap
- Sequencers become a common piece of Lab-kit like RT-PCR machines
- ..Lab scientists will have to learn how to analyse the data!!

# The future-3<sup>rd</sup> gen

- Third-gen sequencing
- Single molecule
- Fast
- Cheap



# Single Molecule

- Sequencing Without amplification
- Real time sequencing
- Less reagent
- Fewer artifacts
- Promise to be:
  - Faster
  - Cheaper
  - Simpler
- As yet no instrument has replaced clonal amplification methods

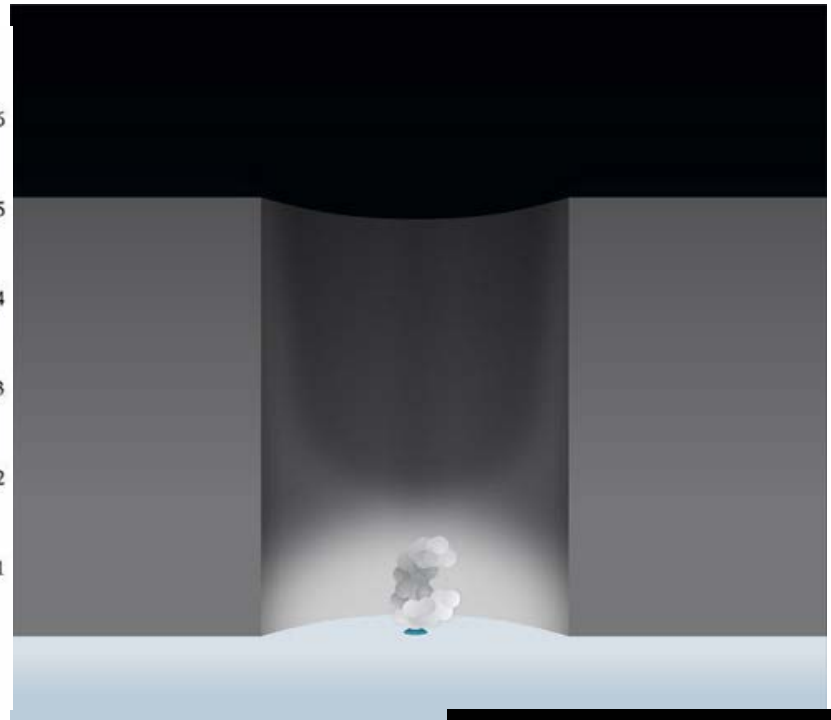
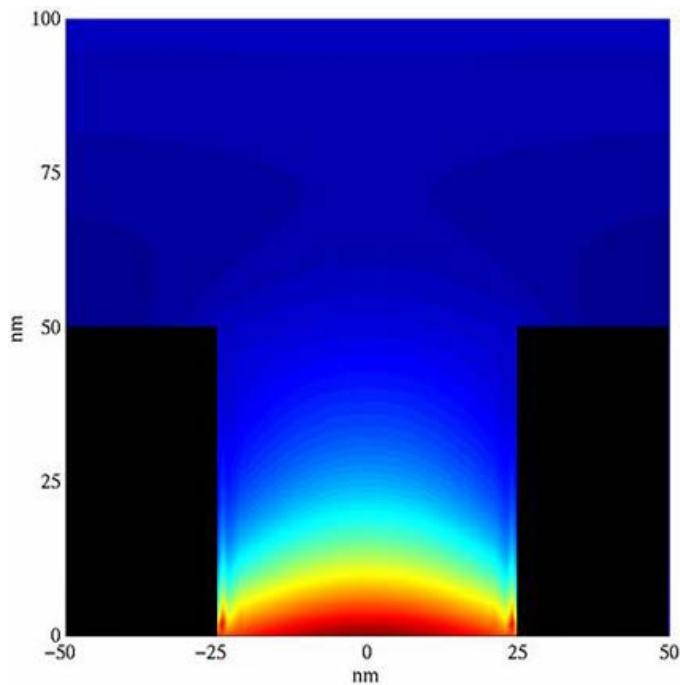


Oxford nanopore

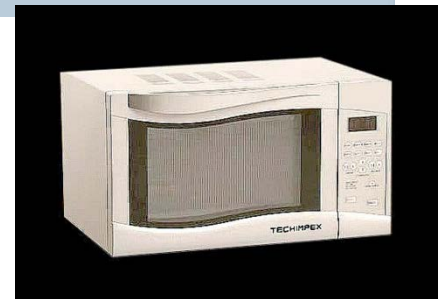


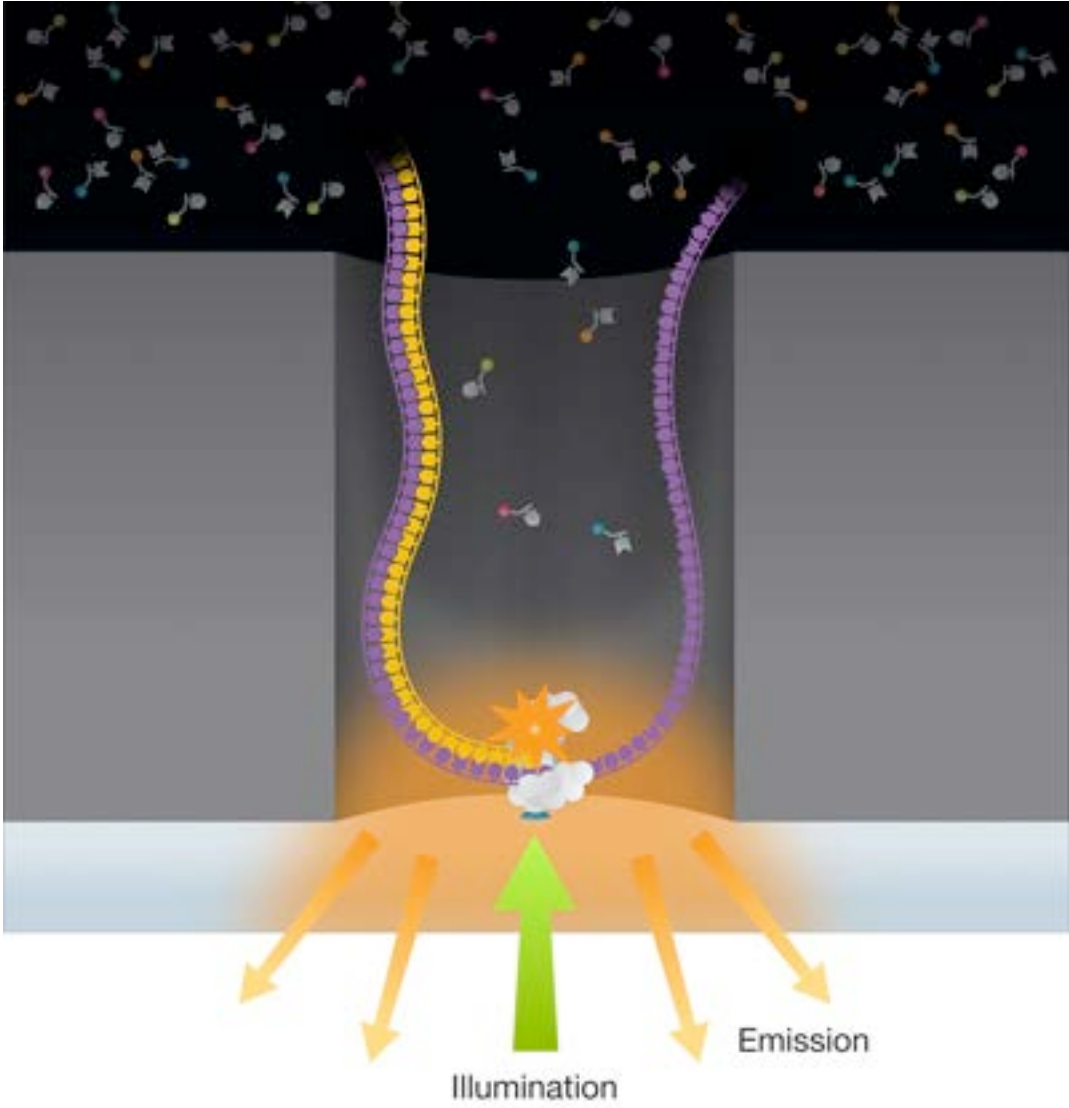
# Pacific Biosciences

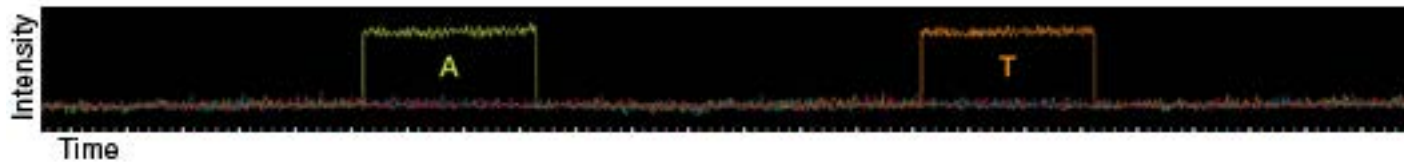
An example of a third generation technology



The zero mode wave  
guide



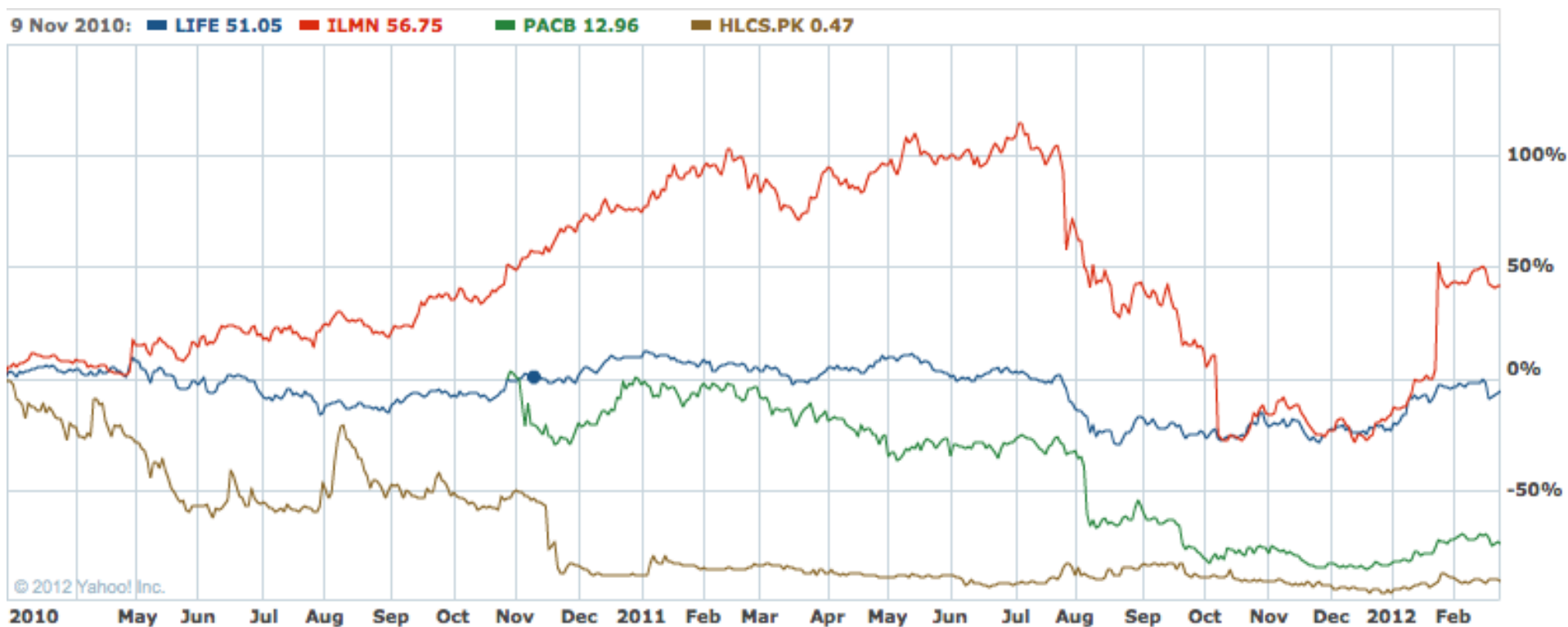




- Long reads > 5kb
- Resequencing of circular molecules for accurate reads.
- Rapid sequencing 30bp per molecule per second
- Single molecules... so very less reagent required
- <http://www.pacificbiosciences.com>
- ...however....
- High error rates
- Low throughput

# So how good is 3<sup>rd</sup> gen

Let the markets decide.....



# The New Next. Next Big Thing: Oxford Nanopore


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## Oxford Nanopore considers flotation within 18 months


Oxford Nanopore, the business that has developed a DNA sequencing device the size of a computer memory stick, could consider a possible \$1bn (£630m) flotation within 18 months.




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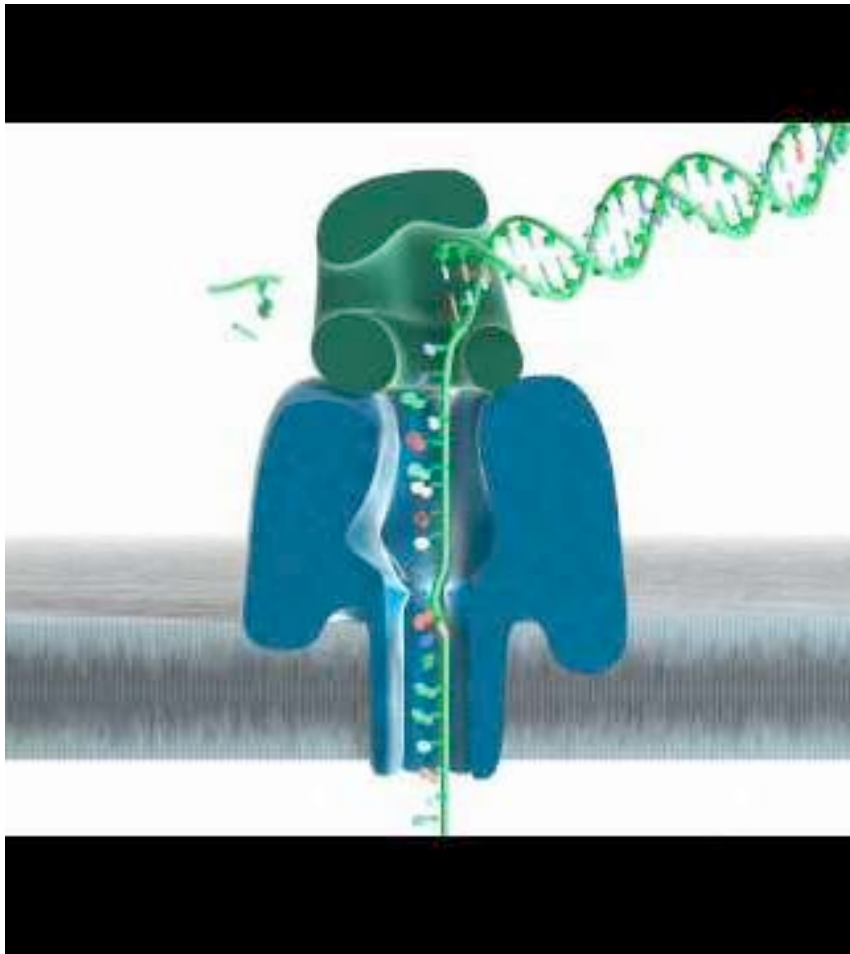
**By Rachel Cooper**  
10:00PM GMT 25 Feb 2012

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 Comment

Spun out of Oxford University in 2005, Oxford Nanopore's shareholders include IP Group, a small-cap business that commercialises discoveries from universities, and Illumina, the American gene-sequencing company.

- Developed sequencing method based of feeding DNA through a biological pore.
- Allows very long reads >10kb
- “Relatively accurate” 96%
- Very cheep devices starting at \$900



- A artificial membrane separates to solutions at different charges.
- A biological pore sits in the membrane
- As DNA is fed through the pore using “ratcheting enzyme”
- The movement of the DNA changes the current between the two
- Each base triplet creates a different electrical signal

# Nanopore sequencing



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platform for real  
time nanopore  
analyses

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GridION

.....potentially very exciting

# Oxford Nanopore:

## Oxford Nanopore valuation doubles following surprise £31.4m fund-raising

WRITTEN BY LAUTARO VARGAS ON 03 MAY 2012. POSTED IN MEDTECH



Oxford Nanopore has announced a surprise £31.4m fundraising, taking the total raised by the Oxford University spin-out since its foundation in 2005 to £105.4m.

The funding is surprising because the company only raised £25m a year ago, stated on its

web site that it wasn't looking for further funding and CEO, Dr Gordon Sanghera, told the Sunday Telegraph just two months ago that the company was well financed for the next stage of growth.

One source told Cabume that the company did not in fact go looking



Sticking with you: Oxford Nanopore's investors pour in another £31.4m to potentially game-changing sequencing tech

- No data has been seen by the community yet
- But ...
- Has already been successful in generating funding!



# Conclusion

- Sequencing will continue to be driven by the clinical investment market
- While markets are convinced that this will generate large revenue there will continue to be well funded development
- So far third generation sequencing has failed to become “mainstream”
- In the short -term we can expect 2<sup>nd</sup> generation short read sequencing to dominate.

- ....questions

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