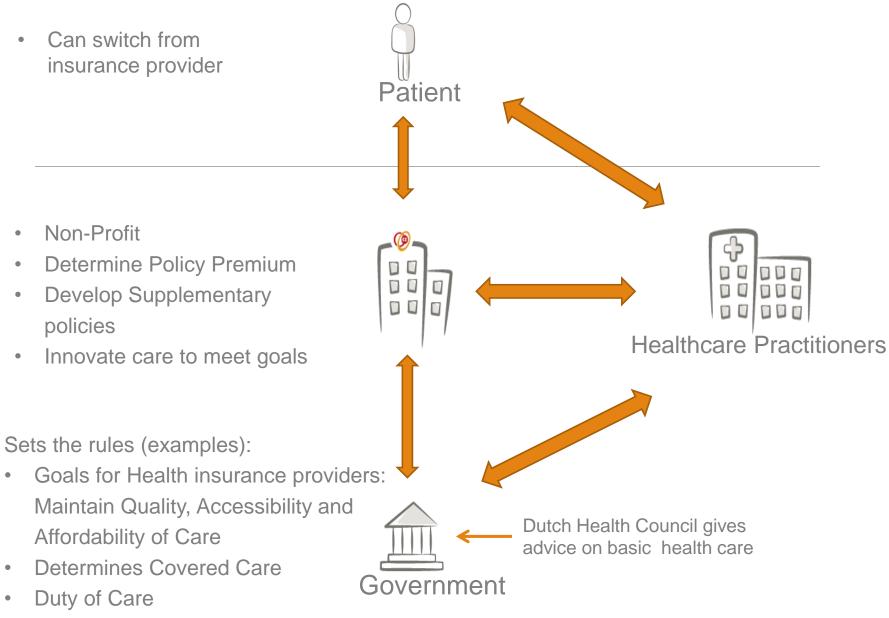


Health Economics of proton therapy from a health insurance perspective

MEDICIS-PROMED Summer School, 7 June 2017 Madelon Johannesma, MSc, PhD _ Health Insurance Company CZ



• Open enrolment obligation

There are a lot of providers The Netherlands....



CZ-group is third largest Dutch health insurer Our Customers



non-profit

Customer satisfaction:	7.9
Policy holders:	3.5 million
Market share:	20%
Premium income:	€ 8.97 billion

Our Operations

Brands: 🧑 OHR	A delta lloyd CZdirect.nl
Healthcare providers contracted:	45.000
Operating costs as % of premium:	3,8%
Solvency II Solvability:	169%
Employees:	2.600

'license to operate': to maintain highquality, accessibility and affordability within the Dutch healthcare system



CZ defines 'the best possible care' as care which creates the most patient value



In any field, improving performance and accountability depends on having a shared goal that unites

Founding father of the concept of 'value' is Michael E. Porter

Value is defined as
Quality
Cost

Health Care Costs are rising exponentially



...and as a consequence the premium

An important cost driver is Health technology

new medical technology is an important determinant in rising health care expenditures 25-75% (average 50%)

	Di Matteo ¹⁵	Jones ¹⁶	Pricewaterhouse Coopers ⁷	Smith et al ¹⁹	Peden and Freeland ¹⁷	Cutler ⁹	Newhouse ¹⁰
Life expectancy/aging	~9%	*	15%**	2%	6%–7%	2%	2%
Administrative costs	*	*	15%***	3%-10%	*	13%	*
Changes in financing	*	*	*	10%	4%5%	10%	10%
Personal income growth	9%-20%	*	*	11%-18%	14%-18%	5%	<23%
Health care prices	*	*	18%	11%-22%	*	19%	*
Technology	~65%	50%-759	625%	38%62%	70%–75%	49%	>65%

Trade-off

A new treatment/technology is considered to be accepted as compared to standard treatment if:

- Better survival rates
- and/or a better QoL
- Acceptable costs

Determined by the Dutch Health Counsil for basic healt care

Current practice: Quick dissiminiation of new technologies without clear proven effectiveness against high costs

e.g. ROBOTIC SURGERY

e.g. PROTON THERAPY





80+ Treatment Rooms to be Operational in Europe by 2020

Number of Treatment Rooms In Each Country

з 3) 18 4 6 HOTTEST! MODERATE JUST HEATING UP 18 Germany 17 Italy 4 Czech Republic Geband € Geband 6 France 3 Sweden 🖀 14 UK 4 Denmark O9 Switzerland 3 Austria 0 2 Belgium 9 Netherlands

Treatment Rooms are Spread Across 35 Proton Therapy Centres* in 12 European Countries: Germany and UK Leading the Pack

Number of Treatment Rooms Per City



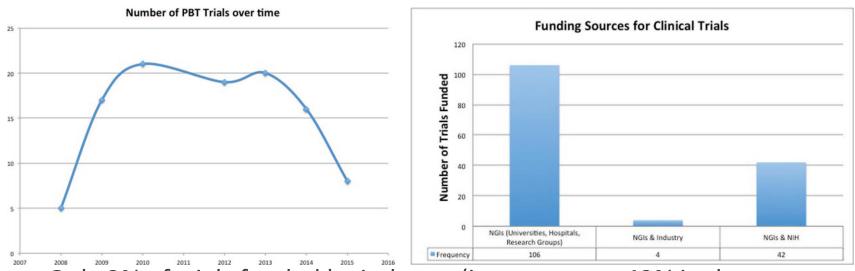
This infographic was created in collaboration with MEDraysintell, in coordination for the forthcoming Proton Therapy Congress hosted by Kisaco Research, taking place in London. September 20-21, 2016. In general.....there are some difficulties in evidence generation of technologies:

As compared to drugs:

- 1. economic evaluation guidelines are written primarily for pharmaceutics
- 2. unlike drugs, medical devices are indivisible, often multiple applications.
- 3. in contrast with drugs, the technical performance of a technology involve interaction with the clinician, while drugs are a classic case of embodied practice
- 4. blinded studies are more difficult to undertake for technologies
- 5. on whom's responsibility? no proof of effectiveness is required for technologies before entry on the market as is with drugs.

focus is on safety and utility, not on effectiveness and costs

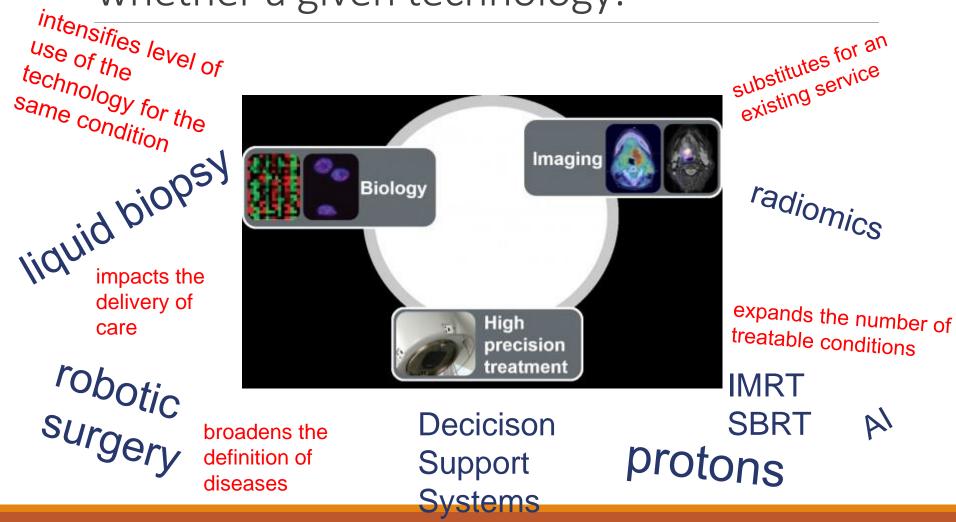
Besides, funding of clinical trials on technologies hampers evidence generation



- Only 3% of trials funded by industry (in contrast to 42% in drugs trials)
- Major funding by universities, hospitals, and research groups (70%)

Evidence gap of technologies can interfere with reimbursement

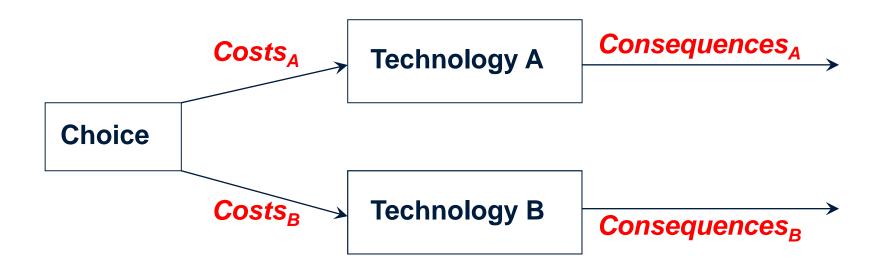
Whether a particular technology increases or decreases costs depends on whether a given technology:



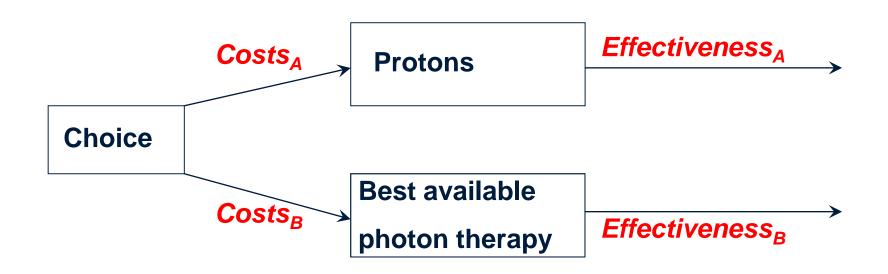
Sorenson et al. ClinicoEconomics and Outcomes Research 2013:5 223-234

To determine the value for money of new technologies cost-effectiveness analyses are performed

 The comparative analysis of alternative courses of action in both their costs and consequences



In case of proton therapy.....



Let's go 10 years back in time.... 2007: no firm conclusion on (cost)effectiveness proton therapy

Conclusion: Existing data do not suggest that the rapid expansion of HT as a major treatment modality would be appropriate. Further research into the clinical and cost-effectiveness of HT is needed. The formation of a European Hadron Therapy Register would offer a straightforward way of accelerating the rate at which we obtain high-quality evidence that could be used in assessing the role of HT in the management of cancer.

Reference:

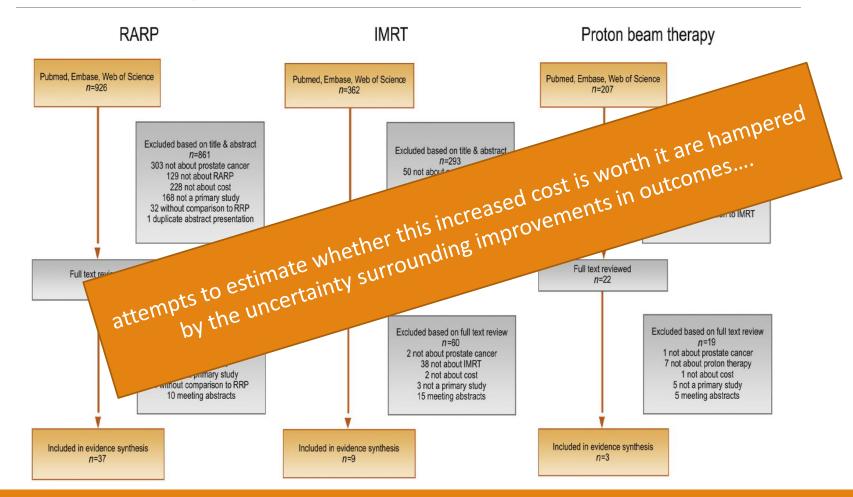
Lodge M, Pijls-Johannesma M, Stirk L, Munro AJ, De Ruysscher D, Jefferson T. A systematic literature review of the clinical and costeffectiveness of hadron therapy in cancer. Radiother Oncol. 2007 May;83(2):110-22.

2013: Limited CEAs for protons for different indications

	Tumor site	Design	Outcome	Result	Report uncertainty
Lundkvist, 2005	Breast	Markov	QALY	€10.130	no
Lundkvist, 2005	Medulloblastoma H&N	Markov	QALY	€10.130	no
Lundkvist, 2005	Prostate	Markov	nany	VDOIN	no
Jakel, 2007	Prostate chorde high level of un assumptions,	suboptimal	method	01093	no
Konski, 2007	assumptions,	Markov	QALY	\$63,578	no
Grutters, 2010	NSCL	Markov	QALY	€67,257	yes
Maboraki, 2010	Rectum	Retrospective analysis	LYG	-	no

Lievens & Pijls-Johannesma, Seminars in Oncology, 2013

2017: Robotic surgery and proton therapy seems more costly than IMRT on prostate cancer treatment.



Schroeck FR, et al. Cost of New Technologies in Prostate Cancer Treatment: Systematic Review of Costs and Cost Effectiveness of Roboticassisted Laparoscopic Prostatectomy, Intensity-modulated Radiotherapy, and Proton Beam Therapy. Eur Urol (2017) So current status (2017): Highly unlikely that protons will be the most cost-effective option for all cancers or even for all patients with a given type of cancer

- Lack of evidence (mainly due to lack of comparable data)
- sub-optimal methodology CEA
- Patient selection needed

Mishra, Mark V. et al. Establishing Evidence-Based Indications for Proton Therapy: An Overview of Current Clinical Trials. Feb 2017;97 (2): 228 – 235

Verma V, Mishra MV, Mehta MP. A systematic review of the cost and cost-effectiveness studies of proton radiotherapy. Cancer. 2016 May 15;122(10):1483-501.

Verma V, Shah C, Rwigema JC, Solberg T, Zhu X, Simone CB 2nd. Cost-comparativeness of proton versus photon therapy. Chin Clin Oncol. 2016 Aug;5(4):56.

Patient selection can be done by modeling, using NTCP & insilico trials

Results of some studies using model-based approaches showed that PBT was not cost-effective for breast-^{1,2} and head-neck cancer³ patients, as it was for a selected group of patients

- **1**. Tommasino F, et al. Model-based approach for quantitative estimates of skin, heart, and lung toxicity risk for left-side photon and proton irradiation after breast-conserving surgery. Acta Oncol. 2017 May;56(5):730-736.
- 2. GY Vega et all. Int J Radiation Oncol Biol Phys, Vol. 95, No. 1, pp. 11-18, 2016
- 3. Ramaekers et al. Protons in head-and-neck cancer: bridging the gap of evidence. Int J Radiat Oncol Biol Phys. 2013 Apr 1;85(5):1282-8.

Example 1

Patient selection by combining in silico data with NTCP

www.redjournal.org

Clinical Investigation

Volume 85 Number 5 2013

Protons in Head-and-Neck Cancer: Bridging the Gap of Evidence

Bram L.T. Ramaekers, MSc, *^{,†} Janneke P.C. Grutters, PhD, * Madelon Pijls-Johannesma, PhD,[†] Philippe Lambin, PhD,[†] Manuela A. Joore, PhD, *^{,‡} and Johannes A. Langendijk, PhD[§]

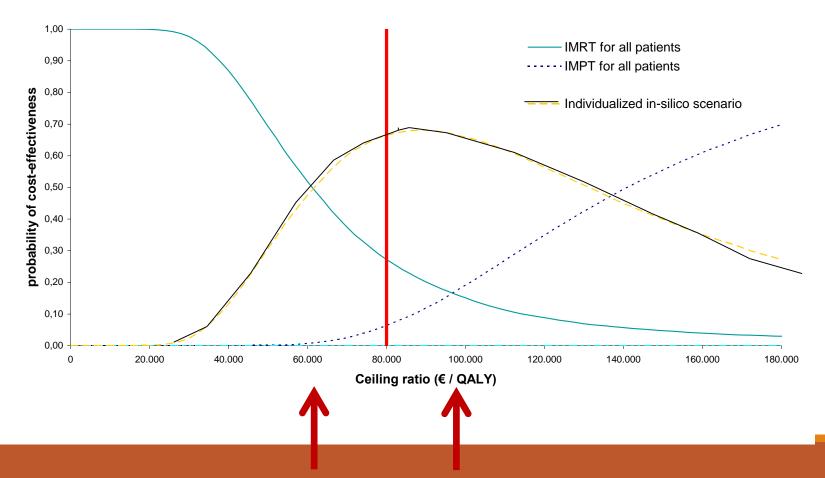
Aim: Given the lack of data, estimate the cost-effectiveness of protons in H&N cancer (IMRT vs IMPT)

Main endpoint: xerostomia and/or dysphagia

- according to available NTCP models*
- dose parameters were derived from in silico trials

IMPT in H&N cancer cost-effective for subgroup of patients





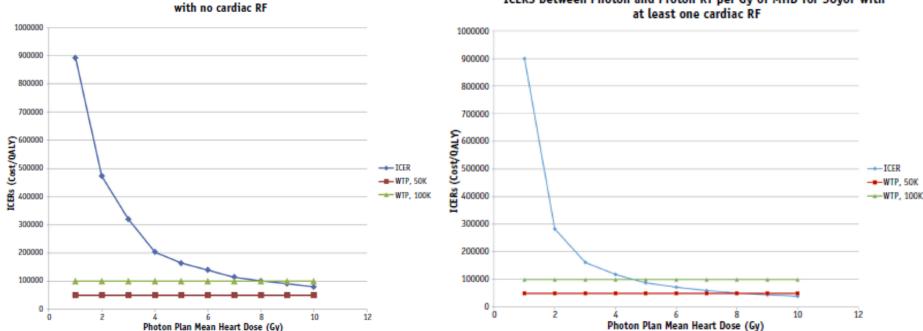
Example 2 Will patient selection contribute to costeffectiveness of protons in breast cancer treatment?

- CEA, Markov model
- photon versus protons
- different strata based on age (40y,50y,60y) and presence/lack of cardiac risk factors (CRF)

Vega et all. Int J Radiation Oncol Biol Phys, Vol. 95, No. 1, pp. 11-18, 2016

Protons for breast-cancer only costeffective if > 1 CRF and MHD > 5 GY

ICERS between Photon and Proton RT per Gy of MHD for 50yoF



ICERS between Photon and Proton RT per Gy of MHD for 50yoF with

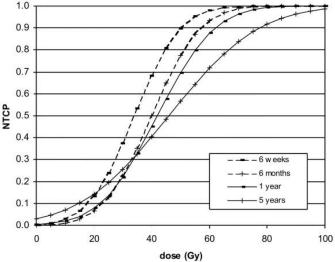
Model-based approach is adopted by the Dutch Health Council

NTCP models will be used to select patients who are likely to benefit from proton therapy (prevention of side effects)

	Radiotherapy and Oncology 107 (2013) 267-273		
	Contents lists available at SciVerse ScienceDirect	Radiotherapy	
	Radiotherapy and Oncology		
FLSEVIER	journal homepage: www.thegreenjournal.com	ATT	
Proton radiotherapy			1

Selection of patients for radiotherapy with protons aiming at reduction of side effects: The model-based approach

Johannes A. Langendijk^{a,*}, Philippe Lambin^b, Dirk De Ruysscher^c, Joachim Widder^a, Mike Bo: Marcel Verheij^e

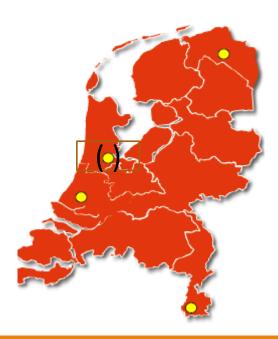


Proton therapy in The Netherlands

Maximum of 4 permits: all granted

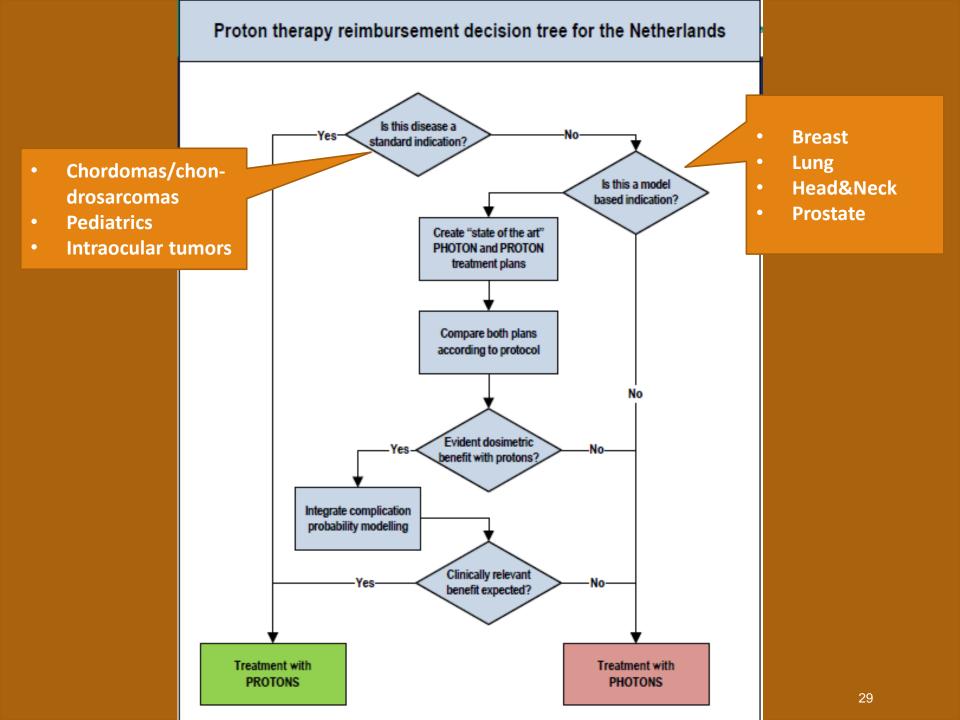
Treatment capacity of 2200 patients/year

- (Amsterdam: 600 pat/yr)
- Groningen: 600 pat/yr
- R'dam/Delft/Leiden: 600 pat/yr
- Maastricht: 400 pat/yr

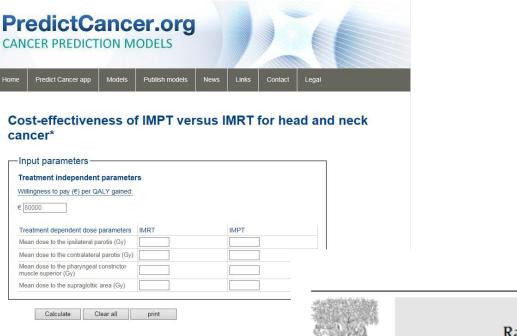


Price agreements made for first 3 years

- Different prices per proton center
- price based on expected number of patients/year
- if less patients will be referred, price per patient will increase
- also start-up costs can be integrated in the price



Decision support systems will play an important role in quickly select the right patient for a certain treatment



	Radiotherapy and Oncology 118 (2016) 281–285	
ie)	Contents lists available at ScienceDirect Radiotherapy and Oncology	Radiotherapy E.Orcology
SEVIER	journal homepage: www.thegreenjournal.com	400

Particle therapy in head and neck cancer

Development and evaluation of an online three-level proton vs photon decision support prototype for head and neck cancer – Comparison of dose, toxicity and cost-effectiveness



Qing Cheng^{a,1}, Erik Roelofs^{a,1}, Bram L.T. Ramaekers^b, Daniëlle Eekers^a, Johan van Soest^a, Tim Lustberg^a, Tim Hendriks^a, Frank Hoebers^a, Hans Paul van der Laan^c, Erik W. Korevaar^c, Andre Dekker^a, Johannes A. Langendijk^c, Philippe Lambin^{a,*}

Summary (1)

It seems hard, if not impossible, to estimate the cost-effectiveness of proton therapy according to basic economic evaluations

- costs are highly variable between countries
- focus is on safety and utility, not on effectiveness and costs
- funding of clinical trials is a problem
- RCT's seams often not possible

Summary (2)

- A model-based approach could be the solution based on subgroup or individual patients.
- Modeling should complement clinical trials, not replace (RCTs remain the ideal tool for research in proton radiotherapy)
- Next, it is possible to assess the effectiveness of proton therapy for individual patients, comparing photon and proton treatments on dose metric, toxicity and cost-effectiveness levels, retrieved from a decision support system.
- Individualized patient selection will enhance the cost-effectiveness of proton therapy (www.predictcancer.org)
- Patient values should be taken into account

Discussion

For a sustainable health care system it is important to control for referring the right patient to the right treatment

- How to standardize and control patient selection?
- Will 2200 patients be referred in The Netherlands, since at current (2016) < 50 patients/year are referred to proton centers abroad?
- So, are we creating overcapacity in The Netherlands?
- What financial consequences will 4 PT centers have for the RT departments /hospitals with only photon therapy?
- Will we succeed to increase value in health care by the introduction of protons in The Netherlands? How can this be measured?

Thanks for your attention!

madelon.johannesma@cz.nl