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# A comparative cost analysis of 25-fraction vs 5-fraction postoperative radiotherapy regimens for breast cancer

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## **Summary**

PURPOSE: To compare the costs of 25-fraction vs 5-fraction postoperative radiotherapy regimens in breast cancer patients. Several clinical trials have confirmed at least comparable safety and efficacy of short hypofractionated partial and whole breast radiation regimens. This study was focused on providing detailed cost data and analysing the advantages or disadvantages of either treatment strategy for patients.

MATERIALS AND METHODS: Calculations were performed based on patient and infra-structure data collected from the department of radiation oncology at Lucerne Cantonal Hospital (LUKS) in Switzerland from 1 July to 31 December 2020. The process maps were created to identify resources used for each radiation therapy option, from initial consultation to treatment completion. Cost comparisons represent the viewpoints of the hospital, insurance coverages and societal costs.

To estimate hospital costs, time-driven activity-based costing was used, including equipment purchase and personnel costs. For insurers, estimates were based on the TARMED tariff system in Switzerland, which is used for billing ambulatory services and reflecting insurance coverage. The social cost was defined as productivity loss (e.g. absence from work) due to treatment appointments.

RESULTS: The 5-fraction regimens resulted in a total of 53% (972 CHF) lower hospital costs, including personnel and equipment resources, a 42% (3153 CHF) reduction of charges to insurers and a 62% (372 CHF) lower social burden in terms of productivity losses due to the patient's absence from work. The major findings from studies using 5-fraction regimens have shown a comparable result in terms of local control and treatment tolerance.

CONCLUSION: A5-fraction radiotherapy regimen in breast cancer patients results in a lower cost than a conventional 25-fraction regimen.

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#### Introduction

Hypofractionated radiation treatment regimens (hRT) involve fewer radiation sessions by using higher doses per session. Hypofractionated radiation treatment is a treat-

ment option for adjuvant radiation therapy in breast cancer patients. Compared to the historical use of hypofractionated regimens, modern technologies allow substantially better sparing of surrounding normal tissues [1]. In addition, radiobiological knowledge has substantially grown based on many clinical trials, empirical observations and radiobiological basic research [1]. In Switzerland, breast cancer is the most common tumour type in women, causing an average of around 1400 deaths per year between 2016 and 2020 [2, 3]. Normofractionated radiation therapy (e.g. 25 × 1.8 or 2.0 Gy per session) with 25 fractions delivered over 5–6 weeks  $\pm$  boost dose application to the tumour bed was the traditional approach for postoperative breast cancer radiation therapy [4-7]. Postoperative whole breast irradiation (WBI) is a worldwide standard after breastconserving tumour surgery to improve locoregional control. Over the last decades, ultra-hypofractionated accelerated partial-breast irradiation (APBI) using e.g. 5-fraction regimens has been prospectively studied for women with early-stage operated breast cancer. The major findings were a high and comparable local control, better treatment tolerance and substantially higher patient comfort compared to whole breast irradiation [8–13]. Similarly, several cohorts and two prospectively randomised trials (FAST and FAST-Forward) [14, 15] using a 5-fraction regimen (± sequential boost) have shown comparable results as for normofractionated radiation therapy also for postoperative whole breast irradiation (e.g.  $5 \times 5.2$  or 5.4 Gy). These ultra-hypofractionated radiation therapy schedules offer the ability to complete whole or partial breast treatment in 1-2 weeks.

At the department of radiation oncology at Lucerne Cantonal Hospital (LUKS) in Switzerland, the clinical implementation of the 5-fraction accelerated partial-breast irradiation course started back in January 2017; the 5-fraction whole breast irradiation schedule according to the FAST-Forward trial was initiated in March 2020.

Despite robust evidence from prospective randomised controlled trials, accelerated partial-breast irradiation and the newer 5-fraction whole breast irradiation are not yet widely used outside the UK regions, due to nuances in local healthcare systems, reimbursement models and professional culture, to name but a few [16, 17].

Meanwhile, the START B radiotherapy schedule using 15 fractions  $\pm$  a boost dose application of 5–10 fractions [18, 19] is generally accepted as a new standard based on superior outcomes compared to normofractionation. In addition to the medical outcome analyses performed previously [5–15, 18], radiation therapy in breast cancer deserves interest for service providers and insurers due to the high costs associated with advanced technology which are burdensome for the hospitals. Nevertheless, the insurance reimbursements (for example, in the Swiss setting) often do not reflect the complexity of the radiation therapy investments [20].

To the best of our knowledge, a comparative economic study for different radiation therapy regimens for breast cancer patients in Switzerland has not been performed so far. The objective of the current analysis is, therefore, to compare a) the costs for the service provider, b) the charges to the insurers and c) the societal cost consequences of 5-fraction versus 25-fraction postoperative radiotherapy regimens for breast cancer.

#### Methods

#### Patient population

Two groups were selected for analysis at the radiation oncology department at Lucerne Cantonal Hospital, Switzerland: first, data from patients treated with the normofractionated whole breast irradiation schedule with 25 fractions (comparison group) as applied until 2015; second, data from patients who underwent the 5-fraction regimens (treatment group) in 1-2 weeks with accelerated partialbreast irradiation or whole breast irradiation (e.g. 5 × 5.2 or 5.4 Gy), initially started in 2017 and fully implemented as a treatment approach in 2020, respectively, to patients aged ≥50 years (accelerated partial-breast irradiation) and >18 years (whole breast irradiation). The approach we are reporting delivers a higher dose of radiation per fraction (treatment session) by Linac-based External Beam Radiotherapy over a shorter overall treatment course compared to conventional fractionation. The major findings from studies using 5-fraction regimens have shown a comparable result in terms of local control and treatment tolerance [8-13].

Patient data related to treatment plans were collected retrospectively via ARIA® software (Varian Medical Systems, Inc.). Follow-up data, staff consultations, infra-structural/processual data and time tracking were applied during the time horizon. All information was retrieved from 1 July to 31 December 2020.

## Model structure and strategies

Time-driven activity-based costing as a bottom-up approach has previously been described by Kaplan and Porter and was defined as a suitable method for cost evaluations of delivering care, including the direct fixed costs of equipment (e.g. purchase price and maintenance) and direct variable costs of labour (e.g. wages and benefits) [21, 22]. The Swiss TARMED tariff was used to determine the charges to insurers for each service provided to patients by the hospital as the provider [23]. The societal cost was defined as a productivity loss due to the absence of patient's work.

To estimate the societal cost, the patients' median monthly gross wages were extracted from the Federal Statistical Office (FSO) for the corresponding years 2015 and 2020. These data were used for female patients aged below 64 years with active professional status and full-time work, living in Switzerland. The estimates provided information on productivity loss due to missed work while attending scheduled treatments at the radiation oncology department. The wages were converted to per-minute values to calculate the cost of time spent in appointments [24] [table 4]. The economic evaluation was carried out according to the Consolidated Health Economic Evaluation Reporting Standards (CHEERS) checklist [25].

To calculate the delivery costs of radiation therapy courses including labour and equipment, the detailed cost-accounting model was developed in Microsoft<sup>©</sup> Excel without the need for any additional statistical tools. To ensure fair time-independent comparisons, costs were discounted by 3% for all cost categories, consistent with recent Swiss economic evaluations [26, 27]. Our cost evaluation analysis includes the following steps:

- Define the care delivery value chain: First, the care delivery value chain is specified to identify the principal activities involved in patient care required to deliver all services in the considered setting, from patient admission to treatment type completion.
- Structure the process map for each care activity: The
  involvement of caregivers for each process was drawn
  chronologically and displayed. The process map function is shown as a flowchart and includes a detailed capacity supplying resources required to patients, from
  initial consultation until treatment delivery (see appendix).
- 3. Time estimates: Measures the time spent by staff and the equipment allocations, estimated as average minutes devoted to each activity throughout the treatment cycle. Staff consultation, engagements from each specialty and observed clinical workflow details was performed to obtain time duration devoted by personnel instead of formal interviews. Note was made of potential outliers of individual factors for patient positioning and treatment activity, which was executed by radiation therapists. Treatment duration was manually recorded by a member of the study and approved by a therapist for a coded sample of 10 treatment plans with the 5-fraction regimen to validate and compare estimates with infra-structural/processual dataset of the 25-fraction regimen.
- 4. Estimate the cost of supplying patient care: The personnel and equipment costs capture both treatment techniques and allow comparisons between two applicable years: 25 fractions performed in 2015 and 5 fractions in 2020. For ethical reasons and to respect privacy, individual salaries of involved personnel was not permitted, but the average salary for the Canton of Lucerne for the relevant years, 2015 and 2020, according to professional title and years of experiences [28]was used for calculations. The equipment costs (e.g. CT scan and linear accelerator) include purchase prices (pre-tax), including VAT (8%) and depreciation for lifetime use of 12 years, and a markup for the yearly cost of maintenance.

- 5. Estimate the capacity and calculate the capacity cost rate: Capacity cost rate was defined as cost per radiation therapy regimen from the first to the last date and expressed in CHF for personnel and equipment. To calculate this, the annual capacity in minutes was estimated for the above resources. The personnel costs were calculated based on annual working days and average minutes spent for the active daily work, adjusted for holidays, weekends and nonpatient-related work (e.g. breaks and meetings). The capacity of equipment was determined based on their clinical availability per year and average minutes of workload per day. For each resource, the annual cost was divided by the annual capacity to determine the capacity cost rate.
- 6. Estimate the total cost of care: Lastly, for each activity from every process, the time estimate was multiplied by the capacity cost rate for all involved resources to calculate the costs. The sum of all activity costs plus the cost of any consumable materials allows for estimating the total cost of patient care in the two radiation therapy regimens, 5-fraction and 25-fraction.

#### 25-fraction radiotherapy

In all patients, a multidisciplinary team made decisions on oncological treatment for newly diagnosed cases, including recommendations for postoperative radiation. Radiation therapy planning and CT simulation were performed postoperatively, and images were imported into the ARIA® radiation therapy planning system (RTPS). Then, radiation oncologists drew radiation therapy planning target volumes, which were then used by dosimetrists and physicians to define the treatment plan, checked by physicists, and finally approved by responsible staff physicians. Linear-based External Beam Radiotherapy delivered the irradiation dose. Lastly, patients were assigned to follow-up visits once a week.

#### 5-fraction radiotherapy

The processing pathway for 5-fraction radiation therapy is similar as for conventional radiation therapy, but with higher daily treatment doses (ultra-hypofractionated) over a shorter duration time (5 workdays) with a total dose of 26–27 Gy. The treatment is delivered by the same linear accelerators using more than two tangential fields, either for whole breast irradiation or accelerated partial-breast irradiation, and deep inspiration breast holds (DIBH) were performed in patients with a left-sided tumour. The patients are assigned two doctor visits to evaluate any potential concerns during the treatment process.

#### Results

#### 25-fraction schedule costs

Tables 1–4 present the results of cost comparisons between two treatment strategies across three domains: hospital costs (e.g. personnel and equipment), insurance expenditures and the economic burden of productivity losses.

## Hospital cost

Tables 1–2 (panel A) show the cost breakdown attributed to the treatment processes including personnel and equipment. Table 1 (panel A) shows the total personnel cost for the 25-fraction regimens which resulted in 1153 CHF per case, and includes MD consultation, radiation oncologist visits, CT scanners, a dosimetrist plan, radiation therapy, medical physicist check and nurse counsel. The most expensive personnel-related activity was attributed to radiation oncology visits (412 CHF), representing 35% of the total cost, followed by the MD's clinical preparation.

Table 2 (panel A) shows the total equipment costs, which resulted in 684 CHF per case, including the CT scan and the linear accelerator (LINAC); the highest cost was for

Table 1:

Cost comparisons: Personnel cost per case. Panel A and B show how the hospital wage expenditure for personnel was distributed per course according to 25-fraction and 5-fraction schedules, and repetitive tasks.

Panel A. Perso	nnel cost for 25-fraction regimen						
Map number	Process	Number of repetitions	Time following care (min)	Number of per- sonnel in- volved	Process cost (CHF/min)	Total cost (CHF)	Total cost (%)
1	MD clinical preparation	5	45	1	1.22	274	23%
2	Radiation oncologist	5	40	1	2.06	412	35%
3	CT scanner	1	15	2	0.67	20	2%
4	Dosimetrist	1	45	1	0.72	32	3%
5	Radiation therapist	25	6.4	2	0.68	217	20%
6	Physicist check	25	6.4	1	0.97	155	13%
7	Nurse counsel	2	15	2	0.72	43	4%
Total						1153	100%
Panel B. Perso	nnel cost for 5-fraction regimen	•					•
Map number	Process	Number of repetitions	Time following care (min)	Number of per- sonnel in- volved	Process cost (CHF/min)	Total cost (CHF)	Total cost (%)
1	MD clinical preparation	2	45	1	1.4	126	21%
2	Radiation oncologist	2	40	1	2.4	192	33%
3	CT scanner	1	15	2	0.8	24	4%
4	Dosimetrist	1	45	1	0.84	37	7%
5	Radiation therapist	5	11	2	0.79	86	15%
6	Physicist check	5	11	1	1.13	62	11%
7	Nurse counsel	2	15	2	0.84	50	9%
Total						577	100%

the linear accelerator (676 CHF), representing 99% of total equipment costs.

#### Insurance expenditures

This relates to the coverage of expenditures incurred by the provision of health services for each patient and recognised by the TARMED tariff with respective tax points, converted into Swiss francs accordingly. Table 3 (panel A) shows the collected tax points for the CT scan, the dosimetrist plan and radiation therapy from the first day until the last treatment. The total charges resulted in 7504 CHF.

## Societal loss

Table 4 (panel A) describes the patient's path and includes doctor consultation, CT scan, check-in, waiting time, radiotherapy and nurse counselling. The total cost due to productivity loss resulted in 601 CHF, based on the gross average income per minute according to FOS data for patients aged between ≥50 years (accelerated partial-breast irradiation) and >18 years (whole breast irradiation) (41.00 CHF/h, in 2015) and estimates the time spent at the radiotherapy unit.

For this analysis, the societal cost included the *productivity loss* due to the patient's absence from work, which was measured according to the average time spent at the radiation oncology centre. The estimate of productivity losses follows the opportunity cost principle, i.e. losses are estimated as the cost of the time spent at the hospital, which

provides implications of loss of workplace performance. The results of time spent at the care unit were multiplied by the number of appointment repetitions and multiplied by the gross wage of each patient group to show the potential impact of productivity loss to their employers.

#### 5-fraction schedule costs

#### Hospital cost

Tables 1–2 (panel B) show the cost breakdown attributed to the treatment processes including personnel and equipment. Table 1 (panel B) shows that the total personnel cost per case for the 5-fraction regimens resulted in 577 CHF, and the process and activities are the same as for the 25-fraction regimen technique, although with fewer repetitive tasks (5 instead of 25 scheduled attendances). The most expensive activity was radiation oncology visits (192 CHF), representing 33% of the total cost. Table 2 (panel B) shows the total equipment cost, which resulted in 288 CHF, including the CT scan and the linear accelerator, the highest cost item (278 CHF) relating to the latter, representing 97% of the total.

## Insurance expenditures

Table 3 (panel B) shows the collected taxpoints for the CT scan, the dosimetrist plan and radiation therapy. The hospital charges resulted in 4351 CHF per treatment course.

Table 2:

Cost evaluation comparison: Equipment cost per case. Panel A and B show the equipment cost for each treatment course, including maintenance for the CT scanner and the linear accelerator (LINAC), based on their cost per year, assuming a lifetime of 12 years and relying on repetitive processes required for radiation therapy regimen use.

Panel A. Equip	ment cost for 25-fraction	n regimen		· · · · · · · · · · · · · · · · · · ·		
Map number	Equipment	Number of repetitions	Duration of session (min)	Cost (CHF/min)	Total cost (CHF)	Total cost (%)
2	CT scan simulation	1	15	0.56	8.4	1%
5	Accelerator machine	25	6.46	4.20	676	99%
Total	'				684	100%
Panel B. Equip	ment cost for 5-fraction	regimen		•		
Map number	Equipment	Number of repetitions	Duration of session (min)	Cost (CHF/min)	Total cost (CHF)	Total cost (%)
2	CT scan simulation	1	15	0.65	9.7	3%
5	Accelerator machine	5	11.4	4.88	278	97%
Total	•				288	100%

Table 3:

Cost evaluation comparison: Insurance cost per case. Panel A and B show the insurance expenditures due to the reimbursement scheme pointing hospital's services according to the TARMED tariff mechanism.

Panel A. Insur	ance cost for the 25-	fraction regimen				
Map number	Process	Taxpoints	Number of repetitions	TARMED (CHF/Tax-points)	Total cost (CHF)	Total cost (%)
2	CT scan	320.58	1	0.72	230	3%
3	Dosimetrist	1338.68	1	0.72	964	13%
5	Rad. therapy	350.61	25	0.72	6310	84%
Total					7504	100%
Panel B. Insur	ance cost for 5-fracti	on regimen				
Map number	Process	Taxpoints	Number of repetitions	TARMED (CHF/Tax- points)	Total cost (CHF)	Total cost (%)
2	CT scan	320.58	1	0.84	269	6%
3	Dosimetrist	1338.68	1	0.84	1124	26%
5	Rad. therapy	704.41	5	0.84	2958	68%
Total					4351	100%

#### Societal loss

Table 4 (panel B) describes the productivity losses due to the patient's path time following treatment schedules. The total cost of the productivity loss due to work absence resulted in 229 CHF. The highest cost element was time spent with doctor consultations (56 CHF), representing 25% of the total and is estimated based on the gross average income per hour according to FOS data for patients aged between ≥50 years (accelerated partial-breast irradiation) and >18 years (whole breast irradiation) (42.00 CHF/h, in 2020) and accounted for time spent at the radiotherapy unit.

#### Cost comparisons

## Hospital cost

From the hospital perspective, a 5-fraction regimen resulted in 50% (576 CHF) lower personnel costs and a 58% (396 CHF) reduction in equipment expenses compared to a standard radiation therapy scheme. By implementing the new treatment technique, in total hospital costs (e.g. personnel + equipment) decreased by 53% (972 CHF) compared to the conventional 25-fraction per case. Furthermore, such a significant hospital cost wage decrease, by 576 CHF when implementing a 5-fraction regimen, due to the reductions of repetitive treatments and activities required by healthcare workers which later results in time savings and reduces hospital gross wage expenditures per treatment course. The radiation therapist (see figure S1 in the appendix) spent less time with the accelerator machine on the 25-fraction (6 minutes) than for the 5-fraction regimen (11 minutes), therefore the process for a fraction resulted in cost variations. However, both treatment regimens come as a package to treat patients in terms of treatment method and reimbursement tariffs.

#### Insurance expenditures

When comparing the payer's expenditures (insurance), the newer treatment technique of a 5-fraction regimen resulted in a decrease of 42% (3153 CHF). The decrease in insurers' spending from 7504 CHF to 4351 CHF was entirely related to the reduction in radiation therapy sessions, from 25 to 5 treatment fractions, and the associated taxpoint tariffs. However, a taxpoint-related surcharge was added for each radiation therapy session in the 5-fraction course because more treatment fields are required compared to the 25-fraction course.

#### Societal loss

The 5-fraction regimen reduced the economic burden by approximately 62% (372 CHF) compared to the 25-fraction regimen. This reduction was attributed to fewer missed workdays for patients assigned to the 5-fraction approach.

#### **Discussion**

This evidence provides a comprehensive analysis of the cost implications when evaluating treatment modalities for breast cancer.

The results show that the 5-fraction regimens resulted in a 53% lower hospital cost, including personnel and equipment resources, a 42% reduction in charges to insurers and a 62% lower social burden in terms of productivity losses due to the patient's absence from work.

The ultra-hypofractionated radiation therapy approach, compared with normo- or mild hypofractionated radiation therapy, offers the benefit to patients of a shorter treatment time as well as less immediate treatment side effects, besides less interference with the patient's normal life activity [29–31]. Porter et al. presented the goal for high-value care with money spent and meeting patients' needs which should be set mutually by physicians and board managers where all actors can benefit, including patients, payers and suppliers [32]. To understand the factual cost and manage a condition, it is crucial to have a methodology that allows cost evaluation not only on a departmental basis but also compares treatment strategies within departments. Researchers from Harvard Business School implemented a

Table 4:
Societal cost: Productivity loss due to the absence of work per case. These tables provide productivity losses in society due to the absence of patient's work due to treatment procedures for the time spent at the hospital. It was accounted for using monthly gross salary (h) according to FOS data respectively.

Panel A. Produ	uctivity loss for 25-fracti	on regimen				
Map number	Process	Number of repetitions	Time following care (min)	Wage loss (CHF/min)	Total wage loss (CHF)	Total loss (%)
1	Doctor consultation	5	40	0.65	130	22%
2	CT scan	1	15	0.65	10	2%
3.1	Check-in	25	5	0.65	82	13%
3.2	Waiting	25	15	0.65	243	40%
4	Rad. therapy	25	6	0.65	97	17%
5	Nurse counsel	2	30	0.65	39	7%
Total productivity loss					601	100%
Panel B. Produ	ectivity loss for 5-fractio	n regimen				
Map number	Process	Number of repetitions	Time following care (min)	Wage loss (CHF/min)	Total wage loss (CHF)	Total loss (%)
1	Doctor consultation	2	40	0.7	56	25%
2	CT scan	1	15	0.7	10.5	5%
3.1	Check-in	5	5	0.7	17.5	8%
3.2	Waiting	5	15	0.7	50.4	22%
4	Rad. therapy	5	15	0.7	52.5	22%
5	Nurse counsel	2	30	0.7	42	18%
Total productiv	vity loss				229	100%

time-driven activity-based costing in several organisations, such as the Schön Klink in Germany and the MD Anderson Cancer Center in the US [22, 32]. The major advantage of using such a bottom-up costing approach for cost calculation in radiation therapy is that it yields more accurate costs and better cost structure; estimating the time in minutes attributed by personnel members and equipment used for breast cancer patients. The major cost drivers at the radiation therapy department at Lucerne Cantonal Hospital are wages, 1153 CHF for a 25-fraction regimen and 577 CHF for a 5-fraction regimen. The major time spent for patients at the radiation therapy department was related to doctor consultations, 3.3 hours for a 25-fraction regimen and 1.32 hours for a 5-fraction regimen, resulting in a productivity loss of 130 CHF and 56 CHF respectively. However, in a more global context, it was well recognised that the radiation therapy costs will depend on the types of technology available for use, i.e. protons are more costly than a photons beam, and the proportion of patients eligible to use [33]. Other studies assessed the use of shortened fractionation radiation therapy approaches and compared them with conventional techniques; the results have shown the cost-effectiveness ratio advantages [34, 35], which are in line with our results.

#### Limitations

The main limitation of this study is its monocentric nature and specific economic evaluation according to the Swiss healthcare system. It represents practices at Lucerne Cantonal Hospital in the Canton of Lucerne, which may limit the generalisability of the findings to other regions.

Despite its advantages on micro-costing accounts, it is clear that the time-driven activity-based costing approach may face challenges when thinking of larger-scale applications other than the radio-oncology units, which may require, for example, more dedicated staff or software solutions that connect budgeting and treatment plan systems. The ultimate goal in cancer care is to define mixed costing models that balance treatment objectives, cost containment and clinical care.

The insurance expenditures decreased significantly by 42% (3153 CHF) when comparing 5-fraction with 25-fraction regimens. However, the TARMED mechanism was not entirely aligned with the full cycle of care coverages in hospital, because it did not address services in terms of collected taxpoints, e.g. some intermediate activities such as doctor visits and nurse consultations were missing to reflect on the TARMED platform for both treatment arms. Furthermore, productivity loss should address follow-up visits and treatment effectiveness after intervention (e.g. work productivity, activities), which was not addressed here due to a lack of available data.

## Conclusion

In conclusion, this study shows that 5-fraction regimens were a less expensive strategy to treat patients than conventional 25-fraction regimens for a full care cycle, the hospital costs, the insurer's expenditures and societal costs. The clinical effectiveness of these two different radiation therapy schemes has been established by several RCTs [10–19]; the economic assessment of the two different ra-

diation therapy schemes reveals that 5-fraction radiation schemes for breast cancer resulted in lower costs for the hospital as a provider and reduced charges for the insurers compared to the conventional 25-fraction regimens. Furthermore, patients treated with the 5-fraction regimen and who are actively working could return to work earlier.

Although 5-fraction radiation protocols generate less revenue, they create an opportunity to treat additional patients, which is crucial for maximising resource utilisation. Our study provides a foundation that can be used to extend analysis to other radiation therapy centres in Switzerland. Considering the increasingly high cost of oncology treatments, cost-benefit analyses are becoming more appreciated for provider investments and reimbursement policy decisions.

## **Data sharing statement**

All data supporting the findings of this study are available within the article and the appendix.

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#### Potential competing interests

All authors have completed and submitted the International Committee of Medical Journal Editors form for disclosure of potential conflicts of interest. No potential conflict of interest related to the content of this manuscript was disclosed.

## References

- Janssen S, Glanzmann C, Lang S, Verlaan S, Streller T, Linsenmeier C, et al. Hypofractionated Radiotherapy for Breast Cancer Including Riskadapted Boost: Update on Tolerance and Efficacy of an Accelerated START A Regime. Anticancer Res. 2016 May;36(5):2513–22.
- Studer G, Collen T, Glanzman C. Radiotherapie bei brusterhaltender Lokalbehandlung des Mammakarzinomes: Teilbrust- statt Ganzbrust-Bestrahlung? Institut für Radio-Onkologie, Kantons-Spital Luzern. Schweiz: LUKS; 2017.
- Office, Federal Statistical. "Cancer, New Cases and Deaths: Numbers, Rates, Median Age and Risk by Cancer Site - 2016-2020 | Table." Federal Statistical Office, December 12, 2023. https://www.bfs.admin.ch/asset/en/29145350
- Trombetta M, Pignol JP, Montemagi P, Brady LW. Alternate Fractionation in Radiotherapy. Springer International Publishing AG; 2018. p. 128. http://dx.doi.org/10.1007/978-3-319-51198-6.
- Frank A Vicini 1, Reena S Cecchini 2, Julia R White 3, Douglas W
  Arthur 4, Thomas B Julian 5, Ra-chel A Rabinovitch 6, Robert R Kuske
  et al. Long-term primary results of accelerated partial breast irradiation
  after breast-conserving surgery for early-stage breast cancer: a randomised, phase 3, equiva-lence trial. Lancet
  2019;394(10215):2155-2164. doi: http://dx.doi.org/10.1016/
  S0140-6736(19)32514-0.
- Effect of radiotherapy after breast-conserving surgery on 10-year recurrence and 15-year breast cancer death: Meta-analysis of individual patient data for 10 801 women in 17 randomised trials. (2011). The Lancet, 378(9804), 1707–1716. https://doi.org/ http://dx.doi.org/10.1016/S0140-6736(11)61629-2.
- Shah C, Vicini F, Shaitelman SF, Hepel J, Keisch M, Arthur D, et al. The American Brachytherapy Society consensus statement for accelerated partial-breast irradiation. Brachytherapy. 2018;17(1):154–70. http://dx.doi.org/10.1016/j.brachy.2017.09.004.

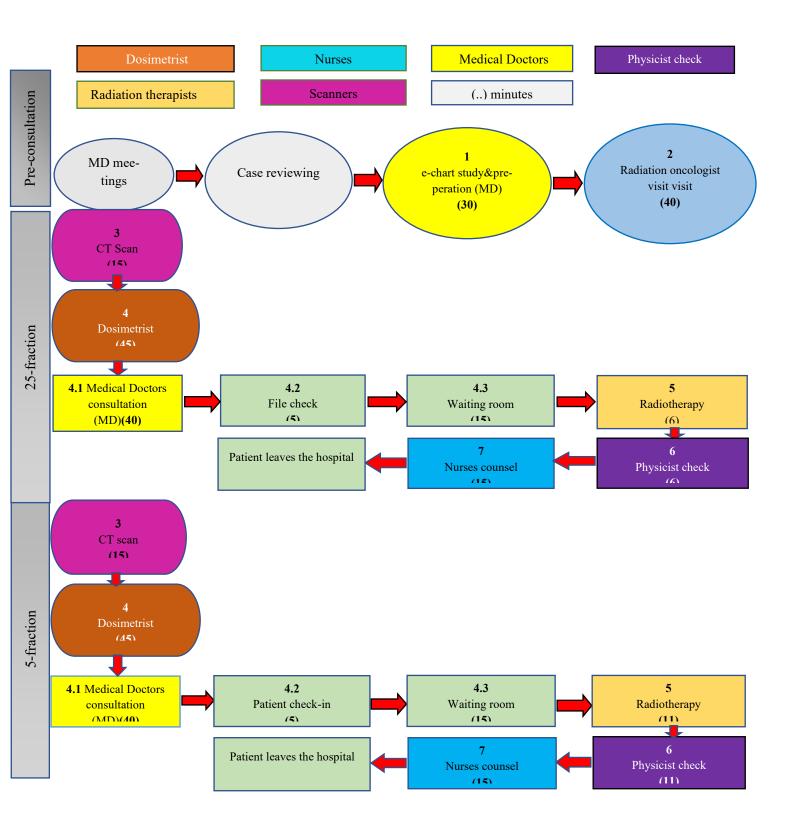
- Moser EC, Vrieling C. Accelerated partial breast irradiation: the need for well-defined patient selection criteria, improved volume definitions, close follow-up and discussion of salvage treatment. Breast. 2012 Dec;21(6):707–15. http://dx.doi.org/10.1016/j.breast.2012.09.014.
- Tann AW, Hatch SS, Joyner MM, Wiederhold LR, Swanson TA. Accelerated partial breast irradiation: Past, present, and future. World J Clin Oncol. 2016 Oct;7(5):370–9. http://dx.doi.org/10.5306/wjco.v7.i5.370.
- Polgár C, Major T, Takácsi-Nagy Z, Fodor J. Breast-Conserving Surgery Followed by Partial or Whole Breast Irradiation: Twenty-Year Results of a Phase 3 Clinical Study. Int J Radiat Oncol Biol Phys. 2021 Mar;109(4):998–1006. http://dx.doi.org/10.1016/ j.ijrobp.2020.11.006.
- Meattini I, Marrazzo L, Saieva C, Desideri I, Scotti V, Simontacchi G, et al. Accelerated Partial-Breast Irradiation Compared With Whole-Breast Irradiation for Early Breast Cancer: Long-Term Results of the Randomized Phase III APBI-IMRT-Florence Trial. J Clin Oncol. 2020 Dec;38(35):4175–83. http://dx.doi.org/10.1200/JCO.20.00650.
- Coles CE, Griffin CL, Kirby AM, Titley J, Agrawal RK, Alhasso A, et al.; IMPORT Trialists. Partial-breast radiotherapy after breast conservation surgery for patients with early breast cancer (UK IMPORT LOW trial): 5-year results from a multicentre, randomised, controlled, phase 3, non-inferiority trial. Lancet. 2017 Sep;390(10099):1048–60. http://dx.doi.org/10.1016/S0140-6736(17)31145-5.
- Whelan TJ, Julian JA, Berrang TS, Kim DH, Germain I, Nichol AM, et al.; RAPID Trial Investigators. External beam accelerated partial breast irradiation versus whole breast irradiation after breast conserving surgery in women with ductal carcinoma in situ and node-negative breast cancer (RAPID): a randomised controlled trial. Lancet. 2019 Dec;394(10215):2165–72. http://dx.doi.org/10.1016/ S0140-6736(19)32515-2.
- Brunt AM, Haviland JS, Sydenham M, Agrawal RK, Algurafi H, Alhasso A, et al. Ten-Year Results of FAST: A Randomized Controlled Trial of 5-Fraction Whole-Breast Radiotherapy for Early Breast Cancer. J Clin Oncol. 2020 Oct;38(28):3261–72. http://dx.doi.org/10.1200/ ICO 19.02750
- Murray Brunt A, Haviland JS, Wheatley DA, Sydenham MA, Alhasso A, Bloomfield DJ, et al.; FAST-Forward Trial Management Group.
   Hypofractionated breast radiotherapy for 1 week versus 3 weeks (FAST-Forward): 5-year efficacy and late normal tissue effects results from a multicentre, non-inferiority, randomised, phase 3 trial. Lancet.
   2020 May;395(10237):1613–26. http://dx.doi.org/10.1016/S0140-6736(20)30932-6.
- Borras JM, Lievens Y, Dunscombe P, Coffey M, Malicki J, Corral J, et al. The optimal utilization proportion of external beam radiotherapy in European countries: an ESTRO-HERO analysis. Radiother Oncol. 2015 Jul;116(1):38–44. http://dx.doi.org/10.1016/j.radonc.2015.04.018.
- Akimoto T, Aoyama H, Chua ML, Jayamanne D, Mizowaki T, Morris L, et al. Challenges and Opportunities With the Use of Hypofractionated Radiation Therapy in Cancer Care: Regional Perspectives From South Korea, Japan, Singapore, and Australia. Adv Radiat Oncol. 2023 Jun;8(6):101291. http://dx.doi.org/10.1016/j.adro.2023.101291.
- 18. Joanne S Haviland, MScJ Roger Owen, FRCRProf John A Dewar, FR-CR Rajiv K Agrawal, FRCRJane Barrett, FRCRProf Peter J Barrett-Lee, MD et al. The UK Standardisation of Breast Radiotherapy (START) trials of radiotherapy hypofractionation for treatment of early breast cancer: 10-year follow-up results of two randomised controlled trials, VOL-UME 14, ISSUE 11, P1086-1094, OCTOBER 2013
- Haviland JS, Mannino M, Griffin C, Porta N, Sydenham M, Bliss JM, et al.; START Trialists' Group. Late normal tissue effects in the arm and shoulder following lymphatic radiotherapy: Results from the UK START (Standardisation of Breast Radiotherapy) trials. Radiother Oncol. 2018 Jan;126(1):155–62. http://dx.doi.org/10.1016/ i.radonc.2017.10.033.
- 20. Picecchi D, Bertram K, Brücher D, Bauer M. Towards novel reimbursement models for expensive advanced therapy medicinal products

- (ATMPs). Swiss Med Wkly. 2020 Dec;150(5153):w20355–20355. http://dx.doi.org/10.4414/smw.2020.20355.
- Schutzer ME, Arthur DW, Anscher MS. Time-Driven Activity-Based Costing: A Comparative Cost Analysis of Whole-Breast Radiotherapy Versus Balloon-Based Brachytherapy in the Management of Early-Stage Breast Cancer. J Oncol Pract. 2016 May;12(5):e584–93. http://dx.doi.org/10.1200/JOP.2015.008441.
- Kaplan RS, Anderson SR. Time-Driven Activity-Based Costing: A Simpler and More Powerful Path to Higher Profits. Harvard Business Press; 2007.
- BAG. B. für G. (n.d.). Tarifsystem TARMED. Retrieved 18 March 2021, from https://www.bag.admin.ch/bag/de/home/versicherungen/krankenversicherung/krankenversicherung-leistungen-tarife/Aerztliche-Leistungen-in-der-Krankenversicherung/Tarifsystem-Tarmed.html
- Office FS. (2022, March 28). Gross monthly wage (median and quartile range) by age, professional position and gender—Private and public sectors combined [T9\_b]—2012, 2014, 2016, 2018, 2020 | Table. Federal Statistical Office. Available from: https://www.bfs.admin.ch/asset/en/21224918
- Husereau D, Drummond M, Augustovski F, de Bekker-Grob E, Briggs AH, Carswell C, et al.; CHEERS 2022 ISPOR Good Research Practices Task Force. Consolidated Health Economic Evaluation Reporting Standards 2022 (CHEERS 2022) Statement: Updated Reporting Guidance for Health Economic Evaluations. Value Health. 2022 Jan;25(1):3–9. http://dx.doi.org/10.1016/j.jval.2021.11.1351.
- Harper S, Kartha M, Mealing S, Borbély YM, Zehetner J. Cost-effectiveness of the RefluxStop device for management of refractory gastroesophageal reflux disease in Switzerland. J Med Econ. 2024;27(1):805–15. http://dx.doi.org/10.1080/13696998.2024.2362564.
- Bommer C, Lupatsch J, Bürki N, Schwenkglenks M. Cost-utility analysis of risk-reducing strategies to prevent breast and ovarian cancer in BRCA-mutation carriers in Switzerland. Eur J Health Econ. 2022 Jul;23(5):807–21. http://dx.doi.org/10.1007/s10198-021-01396-9.
- Amt für Wirtschaft und Arbeit. (n.d.). Kanton Zürich. Retrieved 27
  March 2021. Available from: https://www.zh.ch/de/volkswirtschaftsdirektion/amt-fuer-wirtschaft-und-arbeit.html
- Harnett A. Fewer fractions of adjuvant external beam radiotherapy for early breast cancer are safe and effective and can now be the standard of care. Why the UK's NICE accepts fewer fractions as the standard of care for adjuvant radiotherapy in early breast cancer. Breast. 2010 Jun;19(3):159–62. http://dx.doi.org/10.1016/j.breast.2010.03.029.
- Fletcher GH. Hypofractionation: lessons from complications. Radiother Oncol. 1991 Jan;20(1):10–5. http://dx.doi.org/10.1016/ 0167-8140(91)90106-Q.
- Quirk S, Grendarova P, Craighead P, Phan T, Lesiuk M, Pinilla J, et al. Results of the ACCEL trial: dosimetry in accelerated partial breast irradiation. Radiother Oncol. 2020 Jun;147:50–5. http://dx.doi.org/ 10.1016/j.radonc.2020.03.004.
- Kaplan RS, Porter ME. How to solve the cost crisis in health care. Harv Bus Rev. 2011 Sep;89(9):46–52.
- Lievens Y, Pijls-Johannesma M. Health economic controversy and costeffectiveness of proton therapy. Semin Radiat Oncol. 2013 Apr;23(2):134–41. http://dx.doi.org/10.1016/j.semradonc.2012.11.005.
- Sher, D. J., Wittenberg, E., Suh, W. W., Taghian, A. G., & Punglia, R. S. (2009). Partial-Breast Irradiation Versus Whole-Breast Irradiation for Early-Stage Breast Cancer: A Cost-Effectiveness Analysis. International Journal of Radiation Oncology\*Biology\*Physics, 74(2), 440–446. http://dx.doi.org/10.1016/j.ijrobp.2008.08.015.
- Deshmukh AA, Shirvani SM, Lal L, Swint JM, Cantor SB, Smith BD, et al. Cost-effectiveness Analysis Comparing Conventional, Hypofractionated, and Intraoperative Radiotherapy for Early-Stage Breast Cancer. J Natl Cancer Inst. 2017 Nov;109(11):djx068. http://dx.doi.org/10.1093/ jnci/djx068.

## **Appendix**

## **The Cost Measurement Process**

- 1. Define the care delivery value chain (CDVC): First, the CDVC was specified to identify the principal activities involved in patient care required to deliver all services in the considered setting, from patient admission to treatment type completion.
- 2. Develop process maps of each activity in patient care delivery: Identify the resources involved and any supplies used for the patient.



**FIGURE S1.** Process map illustrates the patient's journey at each activity, with number indicating the sequence of steps and minutes spent on each activity (in brackets). Additionally, colours are used to indicate which specialist or department is responsible for each activity

- 3. Time estimates: Measures the time spent by staff and the equipment allocations, estimated as average minutes devoted to each activity throughout the treatment cycle and activity.
- 4. Estimate the cost of supplying care resources to each patient: In this step, we estimated the direct costs of each resource involved in caring for patients. The personnel and equipment costs capture both treatment techniques, which allow comparisons between two years: 25-fractions performed in 2015 and 5-fractions in 2020.
- 5. Estimate the capacity of each resource and calculate the capacity cost rate (CCR): The CCR is defined as the cost per unit of time (e.g., per minute) for each resource. This is calculated by dividing the annual cost of personnel (e.g., salary, benefits) and similarly for equipment (eannual depreciation, maintenance) by its annual capacity in minutes.

First, estimate the time capacity for each available resource:

*Personnel:* Calculated based on the number of working days, hours per day, and adjusted for nonproductive time (e.g., breaks, training, meetings). For example, when an employee works 50 hours per week, their theoretical capacity is 3,000 minutes (50 hours x 60 minutes).

Equipment: Calculated based on the number of days the equipment is available per year and the hours it can be used each day, adjusted for downtime (e.g., maintenance). For example, when an accelerator machine is available for 249 days per year, and runs 9 hour per day, its annual theoretical capacity is 143,460 minutes (249 days x 9 hours x 60 minutes).

$$CCR(CHF/minute) = \frac{Total\ Annual\ Cost\ of\ the\ Resource}{Total\ Annual\ Capacity\ (in\ minutes)}$$

This formula ensures that the cost of supplying resources is allocated based on the time each resource is actively used, which is a core principle of TDABC.

6. Calculate the total cost of patient care: In this final step, the project team calculates the total cost of treating a patient by multiplying the Capacity Cost Rate (CCR) for each resource used in each task by the time spent on that task. These individual task costs are then summed across all processes involved in the patient's complete cycle of care to determine the total cost of care for the patient.

#### REFERENCES

- 1. Janssen, S., Glanzmann, C., Lang, S., Verlaan, S., Streller, T., Linsenmeier, C., Rades, D., & Studer, G. (2016). Hypofractionated Radiotherapy for Breast Cancer Including Risk-adapted Boost: Update on Tolerance and Efficacy of an Accelerated START A Regime. *Anticancer Research*, *36*(5), 2513–2522.
- 2. Studer, G., Collen, T., Glanzman, C (2017). Radiotherapie bei brusterhaltender Lokalbehandlung des Mammakarzinomes: Teilbrust- statt Ganzbrust-Bestrahlung?. *Institut für Radio-Onkologie, Kantons-Spital Luzern (LUKS), Schweiz.*
- 3. Office, Federal Statistical. "Cancer, New Cases and Deaths: Numbers, Rates, Median Age and Risk by Cancer Site 2016-2020 | Table." Federal Statistical Office, December 12, 2023. https://www.bfs.admin.ch/asset/en/29145350.
- 4. Mark Trombetta, Jean-Pjilippe Pignol, Paolo Montemagi, Luther W. Brady. (2018). Alternate Fractionation in Radiotherapy. Springer International Publishing AG., page 128.
- 5. Frank A Vicini 1, Reena S Cecchini 2, Julia R White 3, Douglas W Arthur 4, Thomas B Julian 5, Ra-chel A Rabinovitch 6, Robert R Kuske et al. Long-term primary results of accelerated partial breast irradiation after breast-conserving surgery for early-stage breast cancer: a randomised, phase 3, equiva-lence trial. Lancet 2019;394(10215):2155-2164. doi: 10.1016/S0140-6736(19)32514-0
- 6. Effect of radiotherapy after breast-conserving surgery on 10-year recurrence and 15-year breast cancer death: Meta-analysis of individual patient data for 10 801 women in 17 randomised trials. (2011). *The Lancet*, 378(9804), 1707–1716. https://doi.org/10.1016/S0140-6736(11)61629-2
- 7. Shah, C., Vicini, F., Shaitelman, S. F., Hepel, J., Keisch, M., Arthur, D., Khan, A. J., Kuske, R., Patel, R., & Wazer, D. E. (2018). The American Brachytherapy Society consensus statement for accelerated partial-breast irradiation. *Brachytherapy*, *17*(1), 154–170. https://doi.org/10.1016/j.brachy.2017.09.004
- 8. Moser EC, Vrieling C. Accelerated partial breast irradiation: the need for well-defined patient selection criteria, improved volume definitions close follow-up and discussion of salvage treatment. Breast. 2012 Dec;21(6):707-15.
- 9. Tann AW, Hatch SS, Joyner MM, Wiederhold LR, Swanson TA. Accelerated partial breast irradiation: Past, present, and future. World J Clin Oncol. 2016 Oct 10;7(5):370-379.
- 10. Polgár C, Major T, Takácsi-Nagy Z, and János Fodor J. Breast-Conserving Surgery Followed by Par-tial or Whole Breast Irradiation: Twenty-Year Results of a Phase 3 Clinical Study. IJROBP, 2021, 109, 4, p998-1006
- 11. Meattini I, Marrazzo L, Saieva C, Desideri I, Scotti V et al. Partial-Breast Irradiation Compared With Whole-Breast Irradiation for Early Breast Cancer: Long-Term Results of the Randomized Phase III APBI-IMRT-Florence Trial. JCO 38, no. 35 (2020) 4175-4183
- 12. Coles, C. E. et al. Partial-breast radiotherapy after breast conservation surgery for patients with early breast cancer (UK IMPORT LOW trial): 5-year results from a multicentre, randomised, controlled, phase 3, non-inferiority trial. Lancet 390, 1048–1060 (2017)
- 13. Whelan, T. J. et al. External beam accelerated partial breast irradiation versus whole breast irradiation after breast conserving surgery in women with ductal carcinoma in situ and node-negative breast cancer (RAPID): a randomised controlled trial. Lancet (2019) doi:10.1016/S0140-6736(19)32515-2
- 14. Brunt, A. M., Haviland, J. S., Sydenham, M., Agrawal, R. K., Algurafi, H., Alhasso, A., Barrett-Lee, P., Bliss, P., Bloomfield, D., Bowen, J., Donovan, E., Goodman, A., Harnett, A., Hogg, M., Kumar, S., Passant, H., Quigley, M., Sherwin, L., Stewart, A., ... Yarnold, J. R. (2020). Ten-Year Results of FAST: A Randomized Controlled Trial of 5-Fraction Whole-Breast Radiotherapy for Early Breast Cancer. *Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 38*(28), 3261–3272. https://doi.org/10.1200/JCO.19.02750
- 15. Murray Brunt, A., Haviland, J. S., Wheatley, D. A., Sydenham, M. A., Alhasso, A., Bloomfield, D. J., Chan, C., Churn, M., Cleator, S., Coles, C. E., Goodman, A., Harnett, A., Hopwood, P., Kirby, A. M., Kirwan, C. C., Morris, C., Nabi, Z., Sawyer, E., Somaiah, N., ... Yarnold, J. (2020). Hypofractionated breast radiotherapy for 1 week versus 3 weeks (FAST-Forward): 5-year efficacy and late normal tissue effects results from a multicentre, non-inferiority, randomised, phase 3 trial. *The Lancet*, 395(10237), 1613–1626. https://doi.org/10.1016/S0140-6736(20)30932-6
- 16. Borras, Josep M., Yolande Lievens, Peter Dunscombe, Mary Coffey, Julian Malicki, Julieta Corral, Chiara Gasparotto, et al. "The Optimal Utilization Proportion of External Beam Radiotherapy in European Countries: An ESTRO-HERO Analysis." *Radiotherapy and Oncology* 116, no. 1 (July 1, 2015): 38–44. https://doi.org/10.1016/j.radonc.2015.04.018.
- 17. Akimoto, Tetsuo, Hidefumi Aoyama, Melvin L.K. Chua, Dasantha Jayamanne, Takashi Mizowaki, Lucinda Morris, Hiroshi Onishi, Si Yeol Song, Youssef H. Zeidan, and Ricky A. Sharma. "Challenges and Opportunities

- With the Use of Hypofractionated Radiation Therapy in Cancer Care: Regional Perspectives From South Korea, Japan, Singapore, and Australia." *Advances in Radiation Oncology* 8, no. 6 (June 14, 2023): 101291. https://doi.org/10.1016/j.adro.2023.101291.
- 18. Joanne S Haviland, MScJ Roger Owen, FRCRProf John A Dewar, FRCR Rajiv K Agrawal, FRCRJane Barrett, FRCRProf Peter J Barrett-Lee, MD et al. The UK Standardisation of Breast Radiotherapy (START) trials of radiotherapy hypofractionation for treatment of early breast cancer: 10-year follow-up results of two randomised controlled trials, VOLUME 14, ISSUE 11, P1086-1094, OCTOBER 2013
- 19. Haviland JS, Mannino M, Griffin C, Porta N, Sydenham M, Bliss JM, et al.; START Trialists' Group. Late normal tissue effects in the arm and shoulder following lymphatic radiotherapy: Results from the UK START (Standardisation of Breast Radiotherapy) trials. Radiother Oncol. 2018 Jan;126(1):155–62 https://doi.org/10.1016/j.radonc.2017.10.033
- 20. Picecchi, Dario, Katrin Bertram, Dominik Brücher, and Michael Bauer. "Towards Novel Reimbursement Models for Expensive Advanced Therapy Medicinal Products (ATMPs)." *Swiss Medical Weekly* 150, no. 5153 (December 30, 2020): w20355–w20355. https://doi.org/10.4414/smw.2020.20355.
- 21. Schutzer, M. E., Arthur, D. W., & Anscher, M. S. (2016). Time-Driven Activity-Based Costing: A Comparative Cost Analysis of Whole-Breast Radiotherapy Versus Balloon-Based Brachytherapy in the Management of Early-Stage Breast Cancer. *Journal of Oncology Practice*, 12(5), e584–e593. https://doi.org/10.1200/JOP.2015.008441
- 22. Kaplan, R. S., & Anderson, S. R. (2007). *Time-Driven Activity-Based Costing: A Simpler and More Powerful Path to Higher Profits*. Harvard Business Press
- 23. BAG, B. für G. (n.d.). *Tarifsystem TARMED*. Retrieved 18 March 2021, from https://www.bag.admin.ch/bag/de/home/versicherungen/krankenversicherung/krankenversicherung-leistungen-tarife/Aerztliche-Leistungen-in-der-Krankenversicherung/Tarifsystem-Tarmed.html
- 24. Office, F. S. (2022, March 28). Gross monthly wage (median and quartile range) by age, professional position and gender—Private and public sectors combined [T9\_b]—2012, 2014, 2016, 2018, 2020 | Table. Federal Statistical Office. https://www.bfs.admin.ch/asset/en/21224918
- 25. Husereau, D., et al., Consolidated Health Economic Evaluation Reporting Standards 2022 (CHEERS 2022) Statement: Updated Reporting Guidance for Health Economic Evaluations. Value Health, 2022. 25(1): p. 3-9.
- 26. Harper, Sam, Muralikrishnan Kartha, Stuart Mealing, Yves M. Borbély, and Jörg Zehetner. "Cost-Effectiveness of the RefluxStop Device for Management of Refractory Gastroesophageal Reflux Disease in Switzerland." *Journal of Medical Economics* 27, no. 1 (December 31, 2024): 805–15. https://doi.org/10.1080/13696998.2024.2362564.
- 27. Bommer, Claudine, Judith Lupatsch, Nicole Bürki, and Matthias Schwenkglenks. "Cost—Utility Analysis of Risk-Reducing Strategies to Prevent Breast and Ovarian Cancer in BRCA-Mutation Carriers in Switzerland." *The European Journal of Health Economics* 23, no. 5 (July 1, 2022): 807–21. https://doi.org/10.1007/s10198-021-01396-9.
- 28. Amt für Wirtschaft und Arbeit. (n.d.). Kanton Zürich. Retrieved 27 March 2021, from;https://www.zh.ch/de/volkswirtschaftsdirektion/amt-fuer-wirtschaft-und-arbeit.html
- 29. Harnett, A. (2010). Fewer fractions of adjuvant external beam radiotherapy for early breast cancer are safe and effective and can now be the standard of care: Why the UK's NICE accepts fewer fractions as the standard of care for adjuvant radiotherapy in early breast cancer. *The Breast*, 19(3), 159–162. https://doi.org/10.1016/j.breast.2010.03.029
- 30. Fletcher, G. H. (1991). Hypofractionation: Lessons from complications. *Radiotherapy and Oncology, 20* (1), 10–15. https://doi.org/10.1016/0167-8140(91)90106-Q
- 31. Quirk, S., Grendarova, P., Craighead, P., Phan, T., Lesiuk, M., Pinilla, J., Liu, H.-W., Wilson, J., Bignell, K., Austin, T., Olivotto, I. A., & Roumeliotis, M. (2020). Results of the ACCEL trial: Dosimetry in accelerated partial breast irradiation. *Radiotherapy and Oncology*, *147*, 50–55. https://doi.org/10.1016/j.radonc.2020.03.004
- 32. Kaplan, R. S., & Porter, M. E. (2011). How to solve the cost crisis in health care. Harv Bus Rev, 89(9), 46-52.
- 33. Lievens, Y., & Pijls-Johannesma, M. (2013). Health Economic Controversy and Cost-Effectiveness of Proton Therapy. *Seminars in Radiation Oncology*, *23*(2), 134–141. https://doi.org/10.1016/j.semradonc.2012.11.005
- 34. Sher, D. J., Wittenberg, E., Suh, W. W., Taghian, A. G., & Punglia, R. S. (2009). Partial-Breast Irradiation Versus Whole-Breast Irradiation for Early-Stage Breast Cancer: A Cost-Effectiveness Analysis. *International Journal of Radiation Oncology\*Biology\*Physics*, 74(2), 440–446.
- 35. Deshmukh, A. A., Shirvani, S. M., Lal, L., Swint, J. M., Cantor, S. B., Smith, B. D., & Likhacheva, A. (2017). Cost-effectiveness Analysis Comparing Conventional, Hypofractionated, and Intraoperative Radiotherapy for Early-Stage Breast Cancer. *JNCI: Journal of the National Cancer Institute*, 109(11), djx068. https://doi.org/10.1093/jnci/djx068

	<b>S1:</b> Cost comparisons: Pers	•	er case.				
Map no	A. Personnel cost for 25-fr	No Repe- tition	Time following care	Person- nel in- volved	Process cost (CHF/Min	Total cost (CHF)	Total cos
1	MD clinical preparation	5	(Min) 45	(No)	1.22	274	23
2	Radiation oncologist	5	40	1	2.06	412	35
3	CT-Scanner	1	15	2	0.67	20	2
4	Dosimetrist	1	45	1	0.72	32	3
<del>-</del> 5	Radiation therapist	25	6.4	2	0.68	217	20
6	Physicist check	25	6.4	1	0.97	155	13
7	Nurse counsel	2	15	2	0.72	43	4
/ Total	Nuise coulisei	2	13	2	0.72	1,153	100
	B. Personnel cost for 5-fra					1,155	100
Map no	Process	No repe- tition	Time following care (Min)	Person- nel in- volved (No)	Process cost (CHF/Min	Total cost (CHF)	Total cos (%)
1	MD clinical preparation	2	45	1	1.4	126	21
2	Radiation oncologist	2	40	1	2.4	192	33
3	CT-Scanner	1	15	2	0.8	24	4
4	Dosimetrist	1	45	1	0.84	37	7
5	Radiation therapist	5	11	2	0.79	86	15
6	Physicist check	5	11	1	1.13	62	11
7	Nurse counsel	2	15	2	0.84	50	9
Total	1					577	100

Panels show how the hospital wage expenditure for personnel was distributed per course accordingly to 25-fraction and 5-fraction schedules, and repetitive tasks.

Table S	Table S2. Cost evaluation comparison: Equipment cost per case.							
Panel A. Equipment cost for 25-fraction								
Map no	Equipment	No repeti- tion	Time ses- sion (Min)	Cost (CHF/Min)	Total cost (CHF)	Total cost (%)		
2	CT-Scan simulation	1	15	0.56	8.4	1		
5	Accelerator machine	25	6.46	4.20	676	99		
Total					684	100		
Panel B	. Equipment cost for 5-fr	action						
Map no	Equipment	No repeti- tion	Time ses- sion (Min)	Cost (CHF/Min)	Total cost (CHF)	Total cost (%)		
2	CT-Scan simulation	1	15	0.65	9.7	3		
5	Accelerator machine	5	11.4	4.88	278	97		
Total					288	100		

Panels show the equipment cost for each treatment course, including maintenance for CT-scan and the linear accelerator (LINAC), based on their cost per year, assuming a lifetime of 12 years and relying on repetitive processes required for RT regimens use.

Table S	3. Cost evaluation co	mparison: Insurar	nce cost per case	2.		
Panel A	. Insurance cost for	25-fraction				
Map no	Process	Taxpoints	No repeti- tion	TARMED (CHF/Tax- points)	Total cost (CHF)	Total cost (%)
2	CT scan	320.58	1	0.72	230	3
3	Dosimetrist	1,338.68	1	0.72	964	13
5	Rad. therapy	350.61	25	0.72	6,310	84
Total					7,504	100
Panel B	. Insurance cost for	5-fraction	•	•		•
Map no	Process	Taxpoints	No repetition	TARMED (CHF/Tax- points)	Total cost (CHF)	Total cost (%)
2	CT-scan	320.58	1	0.84	269	6
3	Dosimetrist	1,338.68	1	0.84	1,124	26
5	Rad. therapy	704.41	5	0.84	2,958	68
Total	•				4,351	100

Panels show the insurance expenditures due to the reimbursement scheme pointing hospital's services according to the TARMED tariff mechanism.

Table 9	64. Society cost: Product	ivity loss due to	the absence of v	work per case.				
Panel A. Productivity loss for 25-fraction								
Map no	Process	No repeti- tion	Time follow- ing care (Min)	Wage loss (CHF/Min)	Total wage loss (CHF)	Total loss (%)		
1	Doctor consultation	5	40	0.65	130	22		
2	CT-scan	1	15	0.65	10	2		
3.1	Check-in	25	5	0.65	82	13		
3.2	Waiting	25	15	0.65	243	40		
4	Rad. therapy	25	6	0.65	97	17		
5	Nurse counsel	2	30	0.65	39	7		
Total productivity loss					601	100		
Panel I	B. Productivity loss for 5	-fraction						
Map no	Process	No repeti- tion	Time follow- ing care (Min)	Wage loss (CHF/Min)	Total wage loss (CHF)	Total loss (%)		
1	Doctor consultation	2	40	0.7	56	25		
2	CT-scan	1	15	0.7	10.5	5		
3.1	Check-in	5	5	0.7	17.5	8		
3.2	Waiting	5	15	0.7	50.4	22		
4	Rad. therapy	5	15	0.7	52.5	22		
5	Nurse counsel	2	30	0.7	42	18		
Total p	roductivity loss				229	100		

These tables provide productivity losses in society due to the absence of patient's work due to treatment procedures for the time spent at the hospital. It was accounted for monthly gross salary (h) according to FOS data respectively.