

# Cost-effectiveness of CTA, MRA and DSA in patients with non-traumatic subarachnoid haemorrhage

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## Abstract

**Objectives** Intra-arterial digital subtraction angiography (DSA), magnetic resonance angiography (MRA) and computed tomographic angiography (CTA) are imaging modalities used for diagnostic work-up of non-traumatic subarachnoid haemorrhage. The aim of our study was to compare the cost-effectiveness of MRA, DSA and CTA in the first year after the bleed.

**Methods** A decision model was used to calculate costs and benefits (in quality-adjusted life-years [QALYs]) that accrued to cohorts of 1,000 patients. Costs and characteristics of diagnostic tests, therapy, patients' quality of life and associated costs were respected. The diagnostic strategy with highest QALYs and lowest costs was considered most cost-effective.

**Results** DSA was the most effective diagnostic option, yielding on average 0.6039 QALYs (95 % CI, 0.5761–0.6327) per patient, followed by CTA 0.5983 QALYs (95 % CI, 0.5704–0.6278) and MRA 0.5947 QALYs (95 % CI, 0.5674–0.6237). Cost was lowest for DSA (39,808 €; 95 % CI, 37,182–42,663), followed by CTA (40,748 €; 95 % CI, 37,937–43,831) and MRA (41,814 €; 95 % CI, 38,730–45,146). A strategy of CTA followed by DSA if CTA was negative or coiling deemed not feasible, was as effective as DSA alone at average costs of 39,767€ (95 % CI, 36,903–42,402).

**Conclusion** A combined strategy of CTA and DSA was found to be the most cost-effective diagnostic approach.

## Main Messages

- We defined a standard model for cost-effectiveness analysis in diagnostic imaging.
- Comparing total 1-year health costs and benefits, CTA is superior to MRA.
- A strategy of combining CTA and DSA was found to be the most cost-effective diagnostic approach.

**Keywords** Intracranial aneurysm · Cost effectiveness · Digital subtraction angiography · Magnetic resonance angiography · Computed tomography angiography

## Introduction

Rupture of an intracranial aneurysm is a major cause of death and disability with an overall incidence of approximately 9 per 100,000 [1]. In the group of patients who survive the initial bleed, the risk of rebleeding without treatment is 1–2 % per day for the first 4 weeks with a 70 % case fatality rate [2, 3].

To prevent rebleeding from ruptured aneurysms, treatment is performed as soon as feasible in order to exclude the aneurysm from the arterial circulation [2, 4, 5]. Treatment options are either neurosurgical clipping via craniotomy or endovascular aneurysm occlusion by detachable platinum coils. The ISAT trial demonstrated better clinical outcome and significant risk reduction for endovascular treatment. Coiling is the preferred option in patients where both treatments are feasible [6–9]. Total treatment costs were found to be slightly higher for surgical clipping compared with endovascular coiling [10–13].

Non-traumatic subarachnoid haemorrhage (SAH) is caused in about 85 % of patients by a ruptured intracranial aneurysm [14]. Diagnostic imaging is required to fulfil two tasks: (1) detect the presence of intracerebral aneurysms; (2)

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determine the most suitable treatment: endovascular coiling or surgical clipping.

DSA is the reference standard for detecting aneurysms and for determining the feasibility of coiling. Nevertheless this invasive and labour-intensive technique is relatively expensive and carries discomforts and potential risks [15–17]. Promising non-invasive and less costly diagnostic alternatives are computed tomographic angiography (CTA) and magnetic resonance angiography (MRA). CTA is easily available and can be rapidly performed in an acute setting. The main benefit of MRA is the fact that no ionising radiation is used.

Although costs and characteristics of diagnostic tests and treatment have been reported, we found no studies determining the optimal diagnostic pathway in patients with suspected ruptured intracranial aneurysms. Our study aimed to compare the cost-effectiveness of MRA, CTA and DSA in these patients. Cost-effectiveness of diagnostic tests in follow-up of coiled aneurysms has previously been evaluated by Schaafsma et al. [18].

## Materials and methods

We previously assessed test characteristics and accuracy of MRA (1.5 T) and CTA in 75 patients with non-traumatic SAH [19]. Patients were recruited consecutively between 2004 and 2006, and for all patients MRA, CTA and DSA had been performed. Informed consent was given and this study protocol was approved by the Institutional Ethical Committee. DSA served as the standard of reference for aneurysm presence. Feasibility of coiling depended on the judgment of the interventionist based on DSA, or the actual suitability during a coiling attempt. For CTA and MRA two experienced neuroradiologists determined the presence of an aneurysm and judged suitability for endovascular treatment using common criteria, like dome-to-neck ratio and absence of branches originating from the base of the aneurysm. The observers were blinded to the results of other imaging examinations. Pooled sensitivity of CTA for the detection of aneurysms was 91.5 % (95 % CI, 85.0–95.5) and specificity was 94.4 % (95 % CI, 79.0–99.0). Pooled sensitivity of MRA for the detection of aneurysms was 95.4 % (95 % CI, 89.8–98.1) and specificity was 83.3 % (95 % CI, 66.5–93.0). Pooled sensitivity of CTA in determining feasibility of endovascular coiling was 71.9 % (95 % CI, 59.0–82.1) and specificity was 75.4 % (95 % CI, 62.0–85.5). Pooled sensitivity of MRA in determining feasibility of endovascular coiling was 60.6 % (95 % CI, 48.2–71.7) and specificity was 81.4 % (95 % CI, 68.7–89.9). DSA as standard of reference was regarded to have a sensitivity and specificity for aneurysm detection and determination of best treatment of 100 %.

Evaluation of costs and benefits: a decision tree

A decision tree (using Excel Software) was developed to determine differences in health benefits and costs in patients with ruptured aneurysms for various diagnostic pathways [20]. For patients with acute non-traumatic subarachnoid haemorrhage, the decision tree delivers different health states due to characteristics of diagnostic imaging and choice of therapy. Health states were based on whether patients were alive and, when they were alive, whether they were well or disabled. Figure 1 shows the decision tree for MRA. A hypothetical cohort of patients passes through the model based on a 1-year period. More specifically, a cohort of patients with non-traumatic subarachnoid haemorrhage is divided over the different pathways in the decision tree, based on a set of probabilities that were derived from literature. This allowed us to synthesise evidence and thereby to evaluate health costs of diagnostic test and treatment as well as related quality of life and associated costs determined by diagnostic decision. A provider (or healthcare) perspective was used, which included only direct healthcare costs.

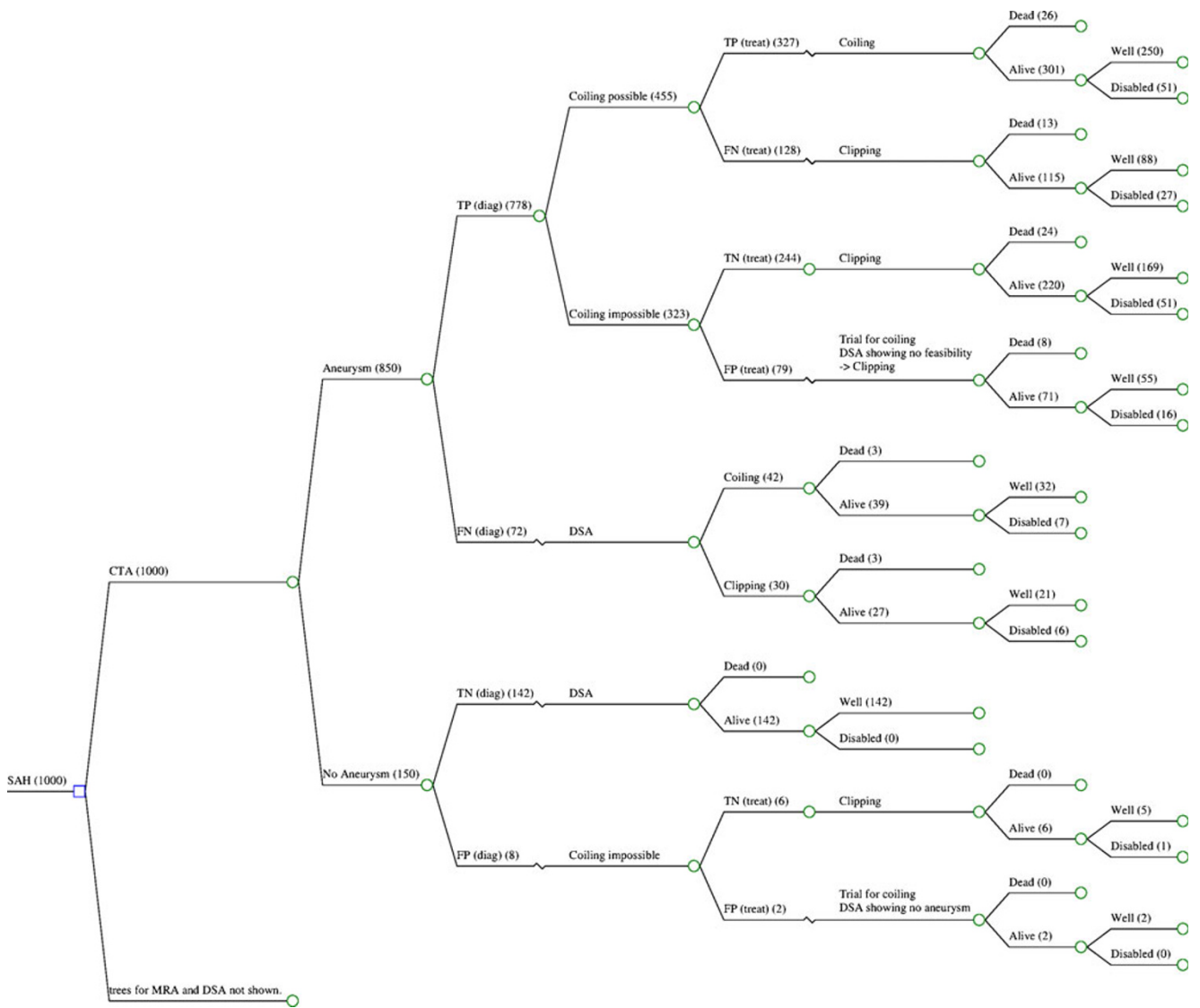
## Basic model and scenario model

In the basic model, patients underwent DSA, MRA or CTA. Following our standard clinical practice, if no aneurysm was detected on CTA or MRA, an additional DSA study was performed. Patients without an intracranial aneurysm were not treated. In patients in whom an aneurysm was detected, feasibility of coiling of the aneurysm was determined. Depending on the result of each test, either a coiling or clipping procedure was performed. In patients where false-positive feasibility of coiling was determined by CTA or MRA, angiography during the coiling procedure would show no feasibility and transfer to surgical clipping would have been performed. In case of false-negative determination of coiling, surgical clipping would have been performed, although coiling would have been feasible.

In a scenario analysis we explored two alternative strategies. CTA and MRA are less expensive and less invasive than DSA. However, they may incorrectly characterise an aneurysm as not suitable for coiling. We therefore analysed whether it is cost-effective to add DSA to the CTA and MRA strategies only if an aneurysm is deemed not suitable for endovascular treatment. In this scenario, in each patient in whom an aneurysm was detected which was deemed not suitable for coiling by MRA or CTA, an additional DSA study was considered to be performed.

## Model parameters

Sensitivity and specificity of the diagnostic tests for detection of aneurysms and determination of treatment possibility were taken as input parameters to the decision model. For the



**Fig. 1** Markov decision tree basic model showing exemplary branch for CTA. Branches for MRA and DSA not shown in detail. *TP* true-positive, *FN* false-negative, *TN* true-negative, *FP* false-positive test

result. Example of absolute patient numbers from base case analysis in *brackets*. Input parameters derived from Table 1

costs of DSA, MRA and CTA, standard prices from the Dutch manual for cost research were used [21]. These include expenditures for personnel, equipment, materials, maintenance, housing, cleaning, administration and overheads. Total 1-year costs of surgical clipping and endovascular coiling were derived from a literature search for western countries [10]. Health outcome after 1 year of treatment was derived from the ISAT trial [6, 7].

Input parameters for related utilities [22] and costs [18, 21, 23], as well as health risk of DSA, are based on available literature [17, 24]. All costs were updated to 2010 by means of national price index figures [25] and expressed in euros (1 €=\$ 1.32). Standard discount rates of 1.5 % for effects and 4 % for costs according to Dutch guidelines were used [21]. Table 1 shows the model input parameters and their sources.

Model assumptions

We assumed that DSA, as the standard of reference, has a sensitivity and specificity of 100 % in detecting aneurysms and determining feasibility of coiling. Furthermore, we assumed no significant gender- or age-related differences in outcome, as well as no influence due to aneurysm size or location. Patients presenting with acute SAH without presence of a ruptured aneurysm were assumed to have no other intracranial vascular pathology to be treated.

Data analysis

In our model, we evaluated the outcome of diagnostic pathways based on the input parameters in Table 1. In the base case

**Table 1** Model input parameters

Model parameter	Mean	SE/SD/range <sup>a</sup>	Distribution	Source
Discount rates				
cDR (cost discount rate)	4 %		fixed	[21]
oDR (outcome discount rate)	1.5 %		fixed	[21]
Treatment independent parameters				
Probabilities, (p)				
pAneurysm in case of non-traumatic SAH	0.85		fixed	[14]
Costs, c (€)				
Diagnostics				
cDSA	725 €		fixed	[21]
cMRA	252 €		fixed	[21]
cCTA	197 €		fixed	[21]
Treatment				
cCoiling	38,238 €	1,833 €	gamma	[10]
cClipping	31,739 €	2,503 €	gamma	[10]
Health state costs				
Number of days in nursing home per year (d)	365			
Costs per day in nursing home (euros/d)	241 €			
cDisability	87,975 €			[21]
Event costs				
cDeath	2,741 €			[18, 23]
Utilities (u)				
uWell	0.78	0.019	beta	[18, 22]
uWell after SAH	0.72	0.65–0.80	triangular	[18, 22]
uDisabled	0.25	0.21–0.30	triangular	[18, 22]
uDead	0.00		fixed	[18, 22]
Clinical outcome probabilities				
Probability of being well after clipping	0.69	0.014	beta	[7]
Probability of being disabled after clipping	0.21	0.013	beta	[7]
pClipdead (probability of being dead after clipping)	0.10	0.009	beta	[7]
pClipwell (probability of being well after survived clipping)	0.77			[7]
Probability of being well after coiling	0.76	0.013	beta	[7]
Probability of being disabled after coiling	0.16	0.011	beta	[7]
pCoildead (probability of being dead after coiling)	0.08	0.008	beta	[7]
pCoilwell (probability of being well after survived coiling)	0.83			[7]
Probability of being well after DSA	0.998	0.001	beta	[17, 24]
Probability of being disabled after DSA	0.002	0.001	beta	[17, 24]
pDSAdead (probability of being dead after DSA)	0.000		fixed	[17, 24]
pDSAwell (probability of being well after survived DSA)	0.998			
Probability of being dead/disabled after surgery without aneurysm				
pSurgdead	0.025	0.004	beta	[30]
pSurgdisab	0.132	0.008	beta	[30]
pCoilTP (probability coiling is feasible in true-positive aneurysm)	0.585	0.061	beta	[19]
pCoilFP (probability coiling is feasible in false-positive aneurysm)	0		fixed	[19]
Treatment dependent parameters				
	Value	SE/SD/range <sup>a</sup>	Distribution	Source
Probabilities				
pMRAtpD (sensitivity of MRA in diagnosing aneurysm)	0.954	0.018	beta	[19]
pMRAtnD (specificity of MRA in diagnosing aneurysm)	0.833	0.061	beta	[19]
pCTAtpD (sensitivity of CTA in diagnosing aneurysm)	0.915	0.024	beta	[19]
pCTAtnD (specificity of CTA in diagnosing aneurysm)	0.944	0.038	beta	[19]

**Table 1** (continued)

pMRAtpT (sensitivity of MRA in determining whether coiling is feasible)	0.606	0.058	beta	<sup>b</sup>
pMRAtnT (specificity of MRA in determining whether coiling is feasible)	0.814	0.050	beta	<sup>b</sup>
pCTAtpT (sensitivity of CTA in determining whether coiling is feasible)	0.719	0.056	beta	<sup>b</sup>
pCTAtnT (specificity of CTA in determining whether coiling is feasible)	0.754	0.057	beta	<sup>b</sup>
pDSAtpD (sensitivity of DSA in diagnosing aneurysm)	1		fixed	[19]
pDSAtnD (Specificity DSA in diagnosing aneurysm)	1		fixed	[19]
pDSAtpT (sensitivity of DSA in determining whether coiling is feasible)	1		fixed	[19]
pDSAtnT (specificity of DSA in determining whether coiling is feasible)	1		fixed	[19]

<sup>a</sup> Range presented for triangular distributions

<sup>b</sup> Data based on study [19]

analysis, the average costs and effects were calculated for a hypothetical cohort of 1,000 patients. We compared total 1-year costs of diagnostic test, treatment option and health state to 1-year health benefits in terms of quality-adjusted life-years (QALYs). The alternative that yielded the highest number of QALYs was considered to be most effective. The current analysis was performed from a decision-analytic perspective. This implies that for an intervention to be adopted, it has to be cost-effective compared with its next best alternative [26]. If a strategy is less costly and more effective, it is superior to and dominates the alternative strategy. If a strategy is more costly and less effective than its alternative, it is dominated by the alternative. In case the strategy is more costly and more effective, or less costly and less effective than the alternative, incremental cost-effectiveness ratios (ICERs) were calculated from each model by dividing the incremental costs by the incremental QALYs. The decision whether the strategy is deemed cost-effective then depends on how much society is willing to pay for a QALY gained. In The Netherlands the informal societal willingness to pay (WTP) threshold level is 80,000 € (\$ 106,000) [27]. If the ICER is lower than this level, we conclude that the strategy is cost-effective compared with the alternative. Additionally, a scenario analysis was performed in which, for all patients, DSA was performed in case an aneurysm was visible but coiling was deemed not feasible.

#### Sensitivity analysis

Uncertainty regarding the model input parameters was explored with a probabilistic sensitivity analysis using Monte Carlo simulation [20, 28]. For this purpose we assigned distributions to all uncertain parameter. With Monte Carlo simulation, 1,000 different samples were taken from these parameter distributions. For each sample, the hypothetical cohort runs through the model based on these sampled probabilities, and costs and effects are derived. This results in 1,000 estimates of costs and effects for the hypothetical cohort, representing the uncertainty in the cost-effectiveness estimation [18]. All assigned distributions are listed in Table 1.

To illustrate the results of the simulation, cost-effectiveness acceptability curves (CEACs) were calculated [29]. CEACs show the probability that a strategy is cost-effective, given different values of willingness to pay for a QALY.

Additionally, univariate sensitivity analyses were performed for test characteristics and costs of DSA, MRA and CTA, as well as for costs of coiling and clipping, to determine association between these model input parameters and cost-effectiveness outcome.

## Results

### Basic model

Comparing expected average 1-year health benefits, DSA was the most effective diagnostic option, yielding 0.6039 QALYs (95 % CI, 0.5761–0.6327), followed by CTA yielding 0.5983 QALYs (95 % CI, 0.5704–0.6278) and MRA yielding 0.5947 QALYs (95 % CI, 0.5674–0.6237). The total expected 1-year health costs were lowest for DSA (39,808 €; 95 % CI, 37,182–42,663), followed by CTA (40,748 €; 95 % CI, 37,937–43,831) and MRA (41,814 €; 95 % CI, 38,730–45,146). Initial diagnostic costs were highest for DSA, while therapy and health state-related costs were lowest for DSA, resulting in DSA being the least costly diagnostic option. CTA as diagnostic test resulted in 128 out of 1,000 patients being referred for clipping although coiling would have been feasible. With MRA, 187 patients out of 1,000 were treated surgically, although endovascular treatment would have been feasible. DSA was both more effective and less costly and dominated CTA and MRA. CTA was more effective and less costly than MRA and thus dominated MRA. Table 2 shows costs and QALYs for the three diagnostic tests.

### Scenario model

The strategy that included DSA for every patient for whom coiling was not deemed feasible on the basis of CTA and MRA, led to equal effectiveness for DSA, MRA and CTA (0.6039 QALYs each). In this approach no patient was

**Table 2** Related 1-year costs and QALYs for MRA, CTA and DSA per patient in the basic model

	MRA		CTA		DSA	
	Costs (€)	QALYs	Costs (€)	QALYs	Costs (€)	QALYs
MRA resp. CTA	252	0.5947	197	0.5983		0.6039
DSA	168 <sup>a</sup>		214 <sup>a</sup>		725	
Total diagnostic	420		411		725	
Treatment	26,734		26,120		25,553	
Death	217		212		205	
Disability costs	14,443		14,005		13,325	
Total	41,814	0.5947	40,748	0.5983	39,808	0.6039

<sup>a</sup> Average costs per patient for DSA performed in case CTA/MRA showing no aneurysm

treated surgically unless coiling was no option. In this scenario, CTA was estimated less costly, reaching a total average 1-year cost per patient of 39,767 €, and therefore cost-effective compared with MRA and DSA, which reached costs of 39,851 € and 39,808 € respectively. Table 3 shows total expected 1-year costs and QALYs as well as incremental costs and QALYs for diagnostic tests in the scenario analysis and basic model.

#### Sensitivity analysis

In univariate sensitivity analyses we explored whether changing individual parameters altered the conclusions of our base case analysis. Assuming equal treatment cost for coiling and clipping did not change the conclusions. Furthermore, results remained stable for the assumption of higher costs of DSA up to factor 2.8. Higher sensitivity and specificity for detection of aneurysms and determination of feasibility of coiling for CTA and MRA up to 96 % or a reduction of sensitivity and specificity for DSA to 90 % also yielded stable results. In the scenario analysis, the strategy where CTA is followed by DSA remained the least costly strategy for a sensitivity of determination of coiling within the range of 60–100 %. The probabilistic sensitivity analysis showed that in the basic model analysis, DSA had a 98–100 % probability of being cost-effective. In the scenario analysis, CTA had the highest probability of being cost-effective (91 %), followed by DSA (9 %) and MRA (0 %). Because in the scenario analysis effectiveness is equal for all strategies, CEACs are constant over the different values of willingness to pay for a QALY.

#### Discussion

When a patient presents with a suspected ruptured intracranial aneurysm, imaging is a challenge. Due to high risk of case fatality in ruptured intracranial aneurysms, timely detection of a causative aneurysm and determination of

appropriate treatment are mandatory. Based on imaging results, patients may be left untreated, or their aneurysms clipped surgically, or they receive endovascular treatment. In recent years, much research has been done to evaluate the best treatment options and to optimise the approach to imaging modalities. In particular, the role and necessity of DSA in this patient population has been widely discussed [31–34]. In our study we compared the diagnostic work-up by CTA, MRA and DSA based on a cost-effectiveness approach. We found that DSA is more cost-effective than CTA, which is in turn superior to MRA. A combination strategy of CTA followed by DSA if endovascular treatment is deemed not feasible was found to be as effective as DSA alone at slightly lower expected costs.

Our analysis is based on a 1-year horizon. This holds potential conflict concerning the long-term validity of the results. Probabilities of health outcome which we clustered from the ISAT trial are relatively stable in the long run and do not change our results [8]. Incomplete occlusion and refilling of the aneurysm is a complication occurring mainly in coiling which can cause rebleeding, resulting in a higher rate of follow-up imaging costs and re-interventions in the endovascular group. The rate of recanalisation is highest in the first months and decreases significantly over time [35–38]. Follow-up for detection of recanalisation after 6 months may not be necessary [36]. Wolstenholme et al. [10] determined cost of coiling and clipping in the first year for western countries. Calculation of total costs for both treatment options included not only cost for material, intensive care and standard unit days and medication but also cost of follow-up imaging and re-intervention for the first year. Taking the results from literature, we can assume that the validity of our data will be stable for the following years.

Our basic model results show that DSA is the most cost-effective imaging modality and is superior to both MRA and CTA. Although initial test costs are about three times higher than those of MRA and CTA, optimal detection of aneurysms and determination of treatment in DSA yield lower overall 1-

**Table 3** Total one-year outcome and incremental cost-effectiveness ratio (ICER) for basic and scenario model

	Costs (€)	(95 % CI)	QALYs	(95 % CI)		
<b>Basic model</b>						
DSA	39,808	(37,182 ; 42,663)	0.6039	(0.5761 ; 0.6327)		
CTA	40,748	(37,937 ; 43,831)	0.5983	(0.5704 ; 0.6278)		
MRA	41,814	(38,730 ; 45,146)	0.5947	(0.5674 ; 0.6237)		
<b>Scenario model</b>						
DSA	39,808	(36,982 ; 42,414)	0.6039	(0.5771 ; 0.6333)		
CTA	39,767	(36,903 ; 42,402)	0.6039	(0.5771 ; 0.6333)		
MRA	39,851	(37,003 ; 42,486)	0.6039	(0.5771 ; 0.6333)		
	Incremental costs (€)	(95 % CI)	Incremental QALYs	(95 % CI)	ICER (€ per QALY)	Comparator
<b>Basic model</b>						
DSA						
CTA	940	(10 ; 2,122)	-0.006	(-0.003 ; -0.009)	Dominated	DSA
MRA	2,007	(604 ; 3,767)	-0.009	(-0.005 ; -0.015)	Dominated	DSA
<b>Scenario model</b>						
DSA						
CTA	-40	(-103 ; 20)	0.000	(0.000 ; 0.000)	Dominant	DSA
MRA	84	(-15 ; 98)	0.000	(0.000 ; 0.000)	Dominated	CTA

CI Confidence Interval (calculated based on the probabilistic sensitivity analysis)

year costs per patient and the highest QALYs. The morbidity of DSA, although low, was included as an input parameter in our model [17, 24]. Nevertheless, we did not consider the potential influence of test-related short-term disutility described by Swan et al. [39]. Depending on the severity grade of clinical presentation, the short-term discomfort undergoing DSA experienced by patients with SAH might be of varying importance and influence to our model might be less compared with in elective imaging.

We assumed DSA to be the standard of reference with sensitivity and specificity for detection and determination of treatment of 100 %. It is debatable whether sensitivity of DSA to cerebral aneurysms is 100 % in an acute setting [40]. When coiling was performed, costs for diagnostic DSA were accounted for separately. Costs of imaging were estimated at 725 €, following the Dutch costing guideline [21]. Actual costs of DSA might be higher than in our model and likely differ in other countries. We therefore performed sensitivity analysis showing stable results for assumption of 90 % sensitivity and specificity in aneurysm detection and coilability, as well as costs up to 2,025 € for DSA. Given this range of diagnostic performance and costs, other researchers or decision-makers can assess transferability of the results to their specific situation and jurisdiction [41]. Differences between United States cost estimates and European cost estimates, in particular, may not change conclusions, as costs are proportionally higher in the United States [42].

The test characteristics for CTA and MRA are based on our study performed with 75 patients. In literature, meta-analyses of test characteristics of CTA as well as other studies show higher sensitivity and specificity for CTA and MRA in the detection of aneurysms and determination of treatment compared with our study [43–45]. Since our basic model results

are stable in sensitivity analysis with assumption of 96 % sensitivity and specificity for aneurysm detection as well as determination of treatment for CTA and MRA, we can assume that our conclusion is valid in a broad range of diagnostic performance. In the scenario model sensitivity analysis, increasing sensitivity and specificity for CTA and MRA shows even increasing preference of our imaging strategy starting with non-invasive imaging compared with only DSA.

Comparing the two non-invasive imaging modalities, CTA dominated MRA in the basic model and thus was superior. This was mainly due to lower sensitivity in determination of feasibility of coiling in MRA. Beside its inferiority in cost-effectiveness, MRA has limitations as first line imaging in acute settings, as it might be unavailable during night hours and not applicable in severe clinical presentation.

Our scenario model with initial CTA followed by DSA in case an aneurysm is detected but deemed not suitable for coiling was found to be the most cost-effective imaging strategy overall. The slightly lower costs compared with only DSA result from the group of patients where on CTA an aneurysm is detected which is deemed feasible for coiling. In this group endovascular treatment will be initiated without previous diagnostic DSA. Nevertheless comparing these two approaches, the average reduction in costs per patient is 40 €, which is marginal in comparison to the overall 1-year costs. In case of suspected SAH, standard diagnostic imaging is non-contrast enhanced CT (NECT), followed by examination of CSF, obtained by lumbar puncture, if NECT is negative [46]. In case a SAH is detected on NECT, performing an additional CTA study can be assumed to be less expensive than the total CTA cost that we used as an input parameter in our model. In fact only additional contrast application would need to be accounted for. Considering this, we might assume that the

actual cost advantage of the scenario strategy will likely be higher.

There are heated discussions about whether or not to always perform DSA in patients with suspected ruptured intracranial aneurysms [47–56]. Our study contributes to this discussion with two important findings. First, DSA is both more effective, yielding higher QALYs, and less costly in overall costs than CTA and MRA. DSA is therefore superior to the non-invasive imaging modalities and should play a key role in diagnostic work-up of non-traumatic SAH. Second, we analysed a scenario that combines invasive and non-invasive diagnostic tests. We found the less costly and therefore most efficient diagnostic approach overall to be a strategy, starting with CTA as the first examination, followed by DSA if coiling is not deemed feasible. This means that every patient undergoes additional DSA unless an aneurysm has been detected and deemed suitable for endovascular coiling. This strategy yields the same amount of QALYs compared with DSA alone in all patients, but results in lower average costs per patient. In comparison to diagnostic DSA, CTA is easily available during night hours and can directly be performed on hospital admission. This further emphasises CTA as a first-line imaging strategy, as it may fasten clinical treatment decisions and therefore help avoid rebleedings.

Besides representing the most cost-effective approach overall, we believe that the strategy of combining CTA and DSA also represents the most practical approach in routine clinical practice. For patients with suspected SAH, we therefore recommend performing a non-contrast enhanced CT scan first. If a SAH is demonstrated, CTA is directly performed. When CTA shows no aneurysm or shows an aneurysm which is not suitable for coiling, an additional DSA study should be performed.

## Conclusions

For diagnostic work-up of suspected ruptured intracranial aneurysms, a combined strategy of CTA followed by DSA if CTA is negative or endovascular treatment deems not feasible is found to be as the most cost-effective approach overall.

**Conflict of interests** None.

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