



Shunting, optic nerve sheath fenestration and dural venous stenting for medically refractory idiopathic intracranial hypertension: systematic review and meta-analysis

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Background: Cerebrospinal fluid (CSF)-diversion procedures have traditionally been the standard of treatment for patients with medically refractive idiopathic intracranial hypertension (IIH). However, dural venous sinus stent (VSS) placement has been described as a safe and effective procedure for the management of medically refractive IIH. We performed a meta-analysis comparing outcomes and complications of CSF-diversion procedures, VSS and optic nerve sheath fenestration (ONSF) for the treatment of medically refractive IIH.

Methods: Electronic searches were performed using six databases from 1988 to January 2017. Data was extracted and meta-analysed from the identified studies.

Results: From 55 pooled studies, there were 538 CSF-diversion cases, 224 dural venous stent placements, and 872 ONSF procedures. Similar improvements were found in terms of postoperative headaches (CSF *vs.* VSS *vs.* ONSF: 84% *vs.* 78% *vs.* 62%, $P=0.223$), papilledema (CSF *vs.* VSS *vs.* ONSF: 71% *vs.* 86% *vs.* 77%, $P=0.192$), whilst visual acuity changes favored venous stenting (CSF *vs.* VSS *vs.* ONSF: 55% *vs.* 69% *vs.* 44%, $P=0.037$). There was a significantly lower rate of subsequent procedures with venous stent placement (CSF *vs.* VSS *vs.* ONSF: 37% *vs.* 13% *vs.* 18%, $P<0.001$), but other complication rates were similar (CSF *vs.* VSS *vs.* ONSF: 13% *vs.* 8% *vs.* 14%, $P=0.28$). Subgroup analysis of lumbar-peritoneal *vs.* ventriculoperitoneal shunts found no differences in symptom improvements, complications and subsequent procedure rates.

Conclusions: Our findings suggest that dural venous sinus stenting may be a viable alternative to traditional surgical interventions in patients who are refractory to medical treatment.

Keywords: Cerebrospinal fluid diversion procedures (CSF-diversion procedures); idiopathic intracranial hypertension (IIH); optic nerve sheath fenestration (ONSF); venous sinus stent placement (VSS)

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Introduction

Idiopathic intracranial hypertension (IIH) is a syndrome defined by increased intracranial pressure without ventriculomegaly or radiographic evidence of a mass lesion, and with normal cerebrospinal fluid (CSF) composition (1). First described by Quincke *et al.* in 1893 as “meningitis serosa”, IIH is a rare condition with only 0.9 cases per 100,000 in the general population (1-4). In comparison to the general population, IIH is most common in obese women aged between 20 and 44 years of age, with an overall prevalence of 15–19/100,000 in North America (2-4).

Given the lack of pathological or radiological evidence, IIH is a diagnosis of exclusion. Currently, IIH is defined by the Modified Dandy Criteria, which for diagnosis, requires signs and symptoms referable only to elevated intracranial pressure, a CSF opening pressure >25 cmH₂O in the lateral decubitus position with normal CSF composition and no evidence of an underlying structural cause on imaging (1). Symptomatically, headache is the presenting complaint in 92–94% of patients (5,6). However, IIH can also lead to papilloedema, which if untreated, can lead to permanent loss of vision (1,7-12). As such, treatment of IIH is of paramount importance.

Despite the pathogenesis of IIH remaining largely speculative, therapeutic developments have substantially advanced patient management (12). Medical management of IIH includes acetazolamide, diuretics, weight loss and serial high-volume lumbar punctures (13). Unfortunately, not all patients respond to the aforementioned medical treatments. In medically refractory IIH, a CSF-diversion procedure, including ventriculoperitoneal (VP) and lumboperitoneal (LP) shunts, and optic nerve sheath fenestration (ONSF) are the next line of treatment (14). Furthermore, cerebral venous sinus stenosis has increasingly been recognised as a cause of increased intracranial pressure. First described by Higgins *et al.* in 2002, dural venous sinus stenting has increasingly been reported to have favourable clinical outcomes in the management of IIH (9,11,15-20).

Traditionally, the treatment for medically refractory IIH has been a CSF-diversion procedure, however alternative techniques have since been developed and used. This article presents an up-to-date meta-analysis and comparison of CSF-diversion procedures, venous sinus stent placement and optic nerve sheath fenestration for the treatment of medically refractory IIH from 1988 to 2017. In order to comprehensively analyse the procedures, an in-depth comparison of all three treatments with respect to post-

operative improvement in headache, papilloedema and visual acuity, complications and the requirement for repeat procedures is completed.

Methods

The present systematic review and meta-analysis was conducted according to recommended guidelines (21,22).

Literature search strategy

Electronic searches were performed using Ovid Medline, PubMed, Cochrane Central Register of Controlled Trials (CCTR), Cochrane Database of Systematic Reviews (CDSR), ACP Journal Club, and Database of Abstracts of Review of Effectiveness (DARE) from their dates of inception to January 2017. To achieve the maximum sensitivity of the search strategy, we combined the terms: “idiopathic intracranial hypertension”, “pseudotumor cerebri”, “benign intracranial hypertension”, “shunt”, “optic nerve sheath decompression”, and “venous stenting”, as either key words or MeSH terms. The reference lists of all retrieved articles were reviewed for further identification of potentially relevant studies and these were then assessed using the inclusion and exclusion criteria.

Selection criteria

Eligible studies for the present systematic review and meta-analysis included those in which patient cohorts underwent an index procedure for treatment of medically refractory IIH via shunting, optic nerve sheath fenestration or venous stenting. Cohorts focusing on patients with repeat procedures were excluded from analysis. Studies that did not include complications as endpoints were excluded. When institutions published duplicate studies with accumulating numbers of patients or increased lengths of follow-up, only the most complete reports were included for quantitative assessment at each time interval. All publications were limited to those involving human subjects and in the English language. Abstracts, case reports, conference presentations, editorials, reviews and expert opinions were excluded.

Data extraction

All data were extracted from article texts, tables and figures. Two investigators independently reviewed each retrieved

article (K Phan, GT Nguyen). Discrepancies between the two reviewers were resolved by discussion and consensus.

Statistical analysis

A meta-analysis of proportions was conducted for the available perioperative and postoperative variables. Heterogeneity was evaluated using Cochran Q and I^2 test. Weighted means were calculated by determining the total number of events divided by total sample size. Subgroup analysis was conducted using mixed-effects meta-regression with a fixed-effect moderator variable for the intervention. All analyses were performed using the “metafor” package for R version 3.02. P values <0.05 were considered statistically significant.

Results

A total of 1,498 references were identified from the electronic database searches. After exclusion and inclusion criteria were applied, a total of 55 studies were included in the analysis (Figure S1), including 22 shunt studies (10,23-43), 21 optic nerve fenestration studies (44-64), and 12 venous stenting studies (9,11,16-20,65-68). Baseline characteristics are summarized in Tables S1-3.

Post-operative improvements in headache, papilloedema and visual acuity, complication rates and the need for a subsequent procedure were analysed for CSF-diversion procedure, venous stent placement and optic nerve sheath fenestration.

CSF-diversion procedure

A total of 22 studies utilizing a CSF-diversion procedure met the inclusion criteria and were analysed (Table S1). This included 538 patients; 85% (398/466) were females. The mean age at presentation was 30.3 years. The mean BMI was 35.2 kg/m² and the mean CSF opening pressure was 41.6 cmH₂O.

The mean follow-up time was 42 months. After the CSF-diversion procedure, 84% of patients had post-operative improvement in headache (95% CI: 0.688–0.923; $I^2=79%$; Figure 1). Seventy-one percent of patients (95% CI: 0.586–0.814; $I^2=51%$; Figure 2) and 55% of patients (95% CI: 0.438–0.654; $I^2=43%$; Figure 3) had post-operative improvement in papilloedema and visual acuity, respectively, following the CSF-diversion procedure.

The complication rate, not including the need for a

subsequent procedure, was 13% (95% CI: 0.082–0.193; $I^2=58%$; Figure S2). The specific complications are listed in Table S1. Thirty-seven percent of patients undergoing a CSF-diversion procedure required a subsequent procedure (95% CI: 0.280–0.476; $I^2=70%$; Figure 4). Specifically, 16 of the 22 studies analysed reported the requirement for a subsequent procedure, with 157 of the 538 patients undergoing an additional 540 procedures.

We also performed a subgroup analysis to compare patients with LP vs. VP shunt methods of CSF-diversion. We found no significant differences between LP vs. VP shunts in terms of improvement in headaches (87.4% vs. 88.2%), papilloedema (77.9% vs. 79%), visual acuity changes (51.8% vs. 49.9%), complication rate (8.1% vs. 15.7%) or subsequent procedure rate (34.5% vs. 47.6%).

Venous stent placement

A total of 12 studies and 224 patients were included in the meta-analysis of dural venous stent placement (Table S2). Eighty-eight percent of patient (197/224) were female and the mean age was 33.4 years. The mean BMI was 34.8 kg/m² and the mean CSF opening pressure was 36.1 cmH₂O.

The mean follow-up time was 20 months. After dural venous stent placement, 78% of patients had post-operative improvement in headache (95% CI: 0.643–0.872; $P<0.01$; Figure 2). Eighty-six percent of patients (95% CI: 0.736–0.930; $P=0.04$; Figure 3) and 69% of patients (95% CI: 0.546–0.801; $P=0.06$; Figure 3) had post-operative improvement in papilloedema and visual acuity, respectively, following dural venous stent placement.

The complication rate, not including the need for a subsequent procedure, was 8% (95% CI: 0.049–0.132; $P=0.71$; Figure S2). Thirteen percent of patients undergoing venous stent placement required a subsequent procedure (95% CI: 0.089–0.186; $P=0.49$; Figure 4). All 12 of the included studies reported the requirement for subsequent procedures, which demonstrated that only 24 of 224 patients underwent additional procedures.

Optic nerve sheath fenestration

A total of 21 studies, including 872 patients and 1,455 eyes, met the inclusion criteria for the meta-analysis (Table S3). Forty-three percent of patients had a unilateral procedure while the remaining 57% of patients underwent bilateral ONSF. Eighty-three percent of patients (522/626) were females and the mean age was 32.2 years. The mean BMI

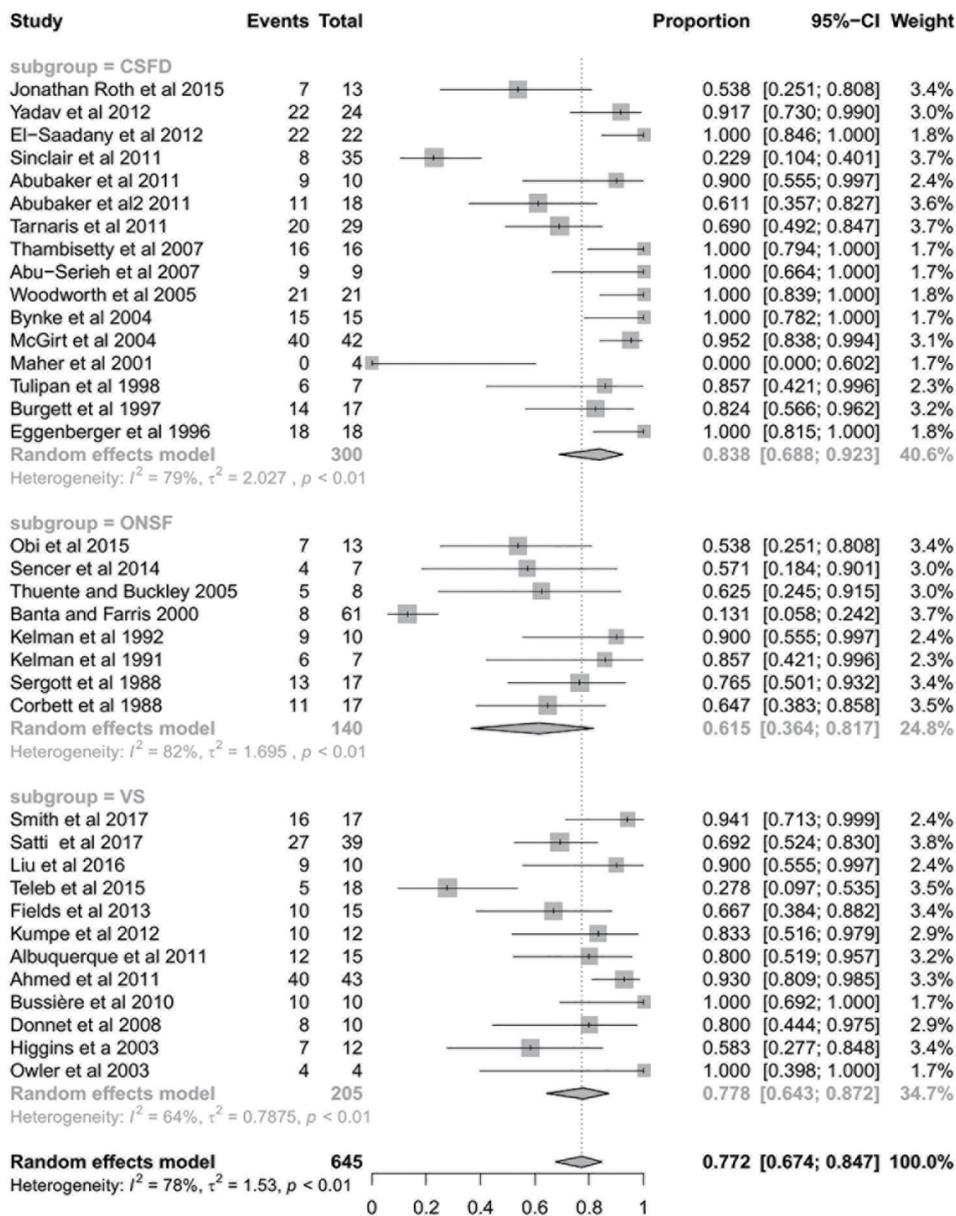


Figure 1 Forest plot of pooled postoperative headache improvement subgrouped according to CSFD, ONSF and VS. CSFD, cerebrospinal fluid diversion; ONSF, optic nerve sheath fenestration; VS, venous stenting.

was 33.5 kg/m² and the mean CSF opening pressure was 33.2 cmH₂O.

The mean follow-up time was 22 months. Following optic nerve sheath fenestration, 62% of patients had post-operative improvement in headache (95% CI: 0.364–0.817; $P < 0.01$; *Figure 1*). Seventy-seven percent of patients (95% CI: 0.598–0.881; $P < 0.01$; *Figure 2*) and 44% of patients (95% CI: 0.320–0.564; $P < 0.01$; *Figure 3*) had post-operative

improvement in papilloedema and visual acuity, respectively, following ONSF.

The complication rate, not including the need for a subsequent procedure, was 14% (95% CI: 0.080–0.223; $P < 0.01$; *Figure S2*). Eighteen percent of patients undergoing ONSF required a subsequent procedure (95% CI: 0.108–0.283; $P < 0.01$; *Figure 4*). Specifically, 16 of 21 studies reported the requirement for subsequent procedures with 111 patients

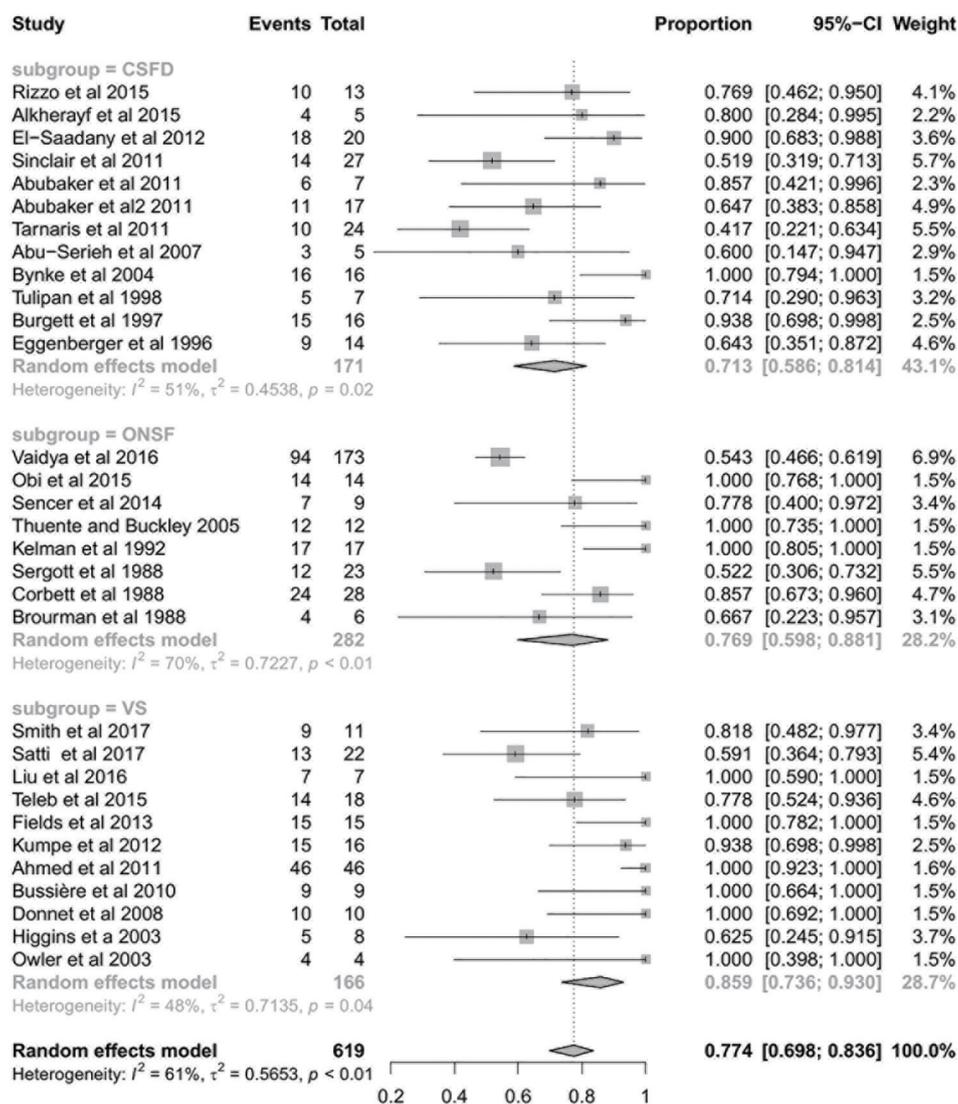


Figure 2 Forest plot of pooled postoperative papilloedema improvement subgrouped according to CSFD, ONSF and VS. CSFD, cerebrospinal fluid diversion; ONSF, optic nerve sheath fenestration; VS, venous stenting.

requiring subsequent procedures out of 699 patients.

Comparison between treatment modalities for medically refractive IIIH

When comparing CSF-diversion procedures, venous stent placement and ONSF with respect to improvement in headache and papilloedema post-operatively, the outcomes were not significantly different between the treatment modalities ($P=0.223$ and 0.192 , respectively). However, post-operative improvement in visual acuity significantly favoured venous stent placement ($P=0.037$).

With respect to complication rates, no statistically significant differences were identified between the three treatment modalities ($P=0.28$), however, there was a significantly lower rate of subsequent procedures required following venous stent placement when compared to CSF-diversion procedures and ONSF ($P<0.001$). Leave-one-out sensitivity analysis did not significantly change the trend of the results.

Discussion

Characterised by increased intracranial pressure without

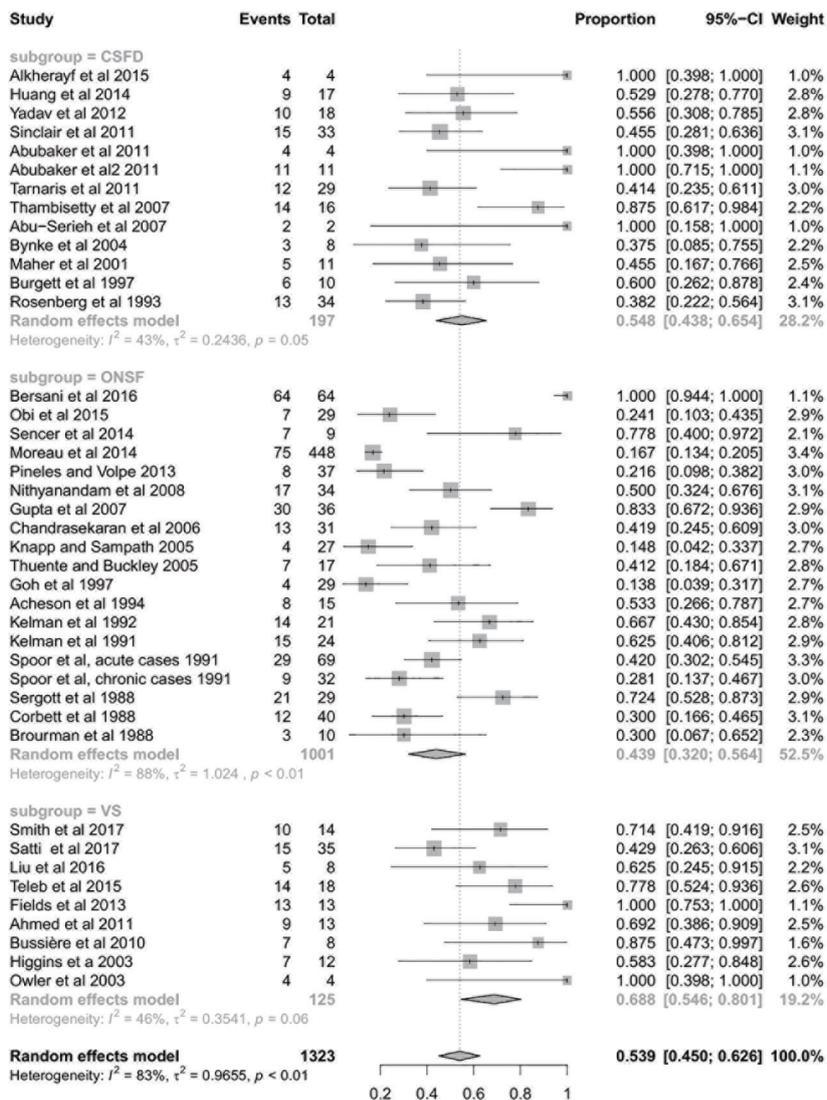


Figure 3 Forest plot of pooled postoperative visual acuity change subgrouped according to CSFD, ONSF and VS. CSFD, cerebrospinal fluid diversion; ONSF, optic nerve sheath fenestration; VS, venous stenting.

a mass lesion or hydrocephalus, IIH classically presents with headache in obese women of childbearing age. Ophthalmologic signs, including diminished visual acuity and papilloedema on fundoscopic examination, frequently present alongside the headache (69). Given that papilloedema associated visual loss is a principle morbidity associated with the condition, the terms “benign intracranial hypertension” and “pseudotumour cerebri” no longer represent current nomenclature and in 2011, the term “idiopathic intracranial hypertension” was adopted (12,69).

Conservative therapy, including weight loss, repeated

high-volume lumbar punctures and medications to reduce CSF production, such as acetazolamide, are the mainstays of treatment. However, some patients are non-responsive to medical therapy and as such, experience progressive worsening of symptoms or develop visual changes. For patients with medically refractive IIH, these patients are traditionally referred for ONSF or a CSF-diversion procedure including VP and LP shunting (70,71). However, more recently, dural venous sinus stent placement has been described in the literature as a safe and effective procedure for the management of medically refractive IIH.

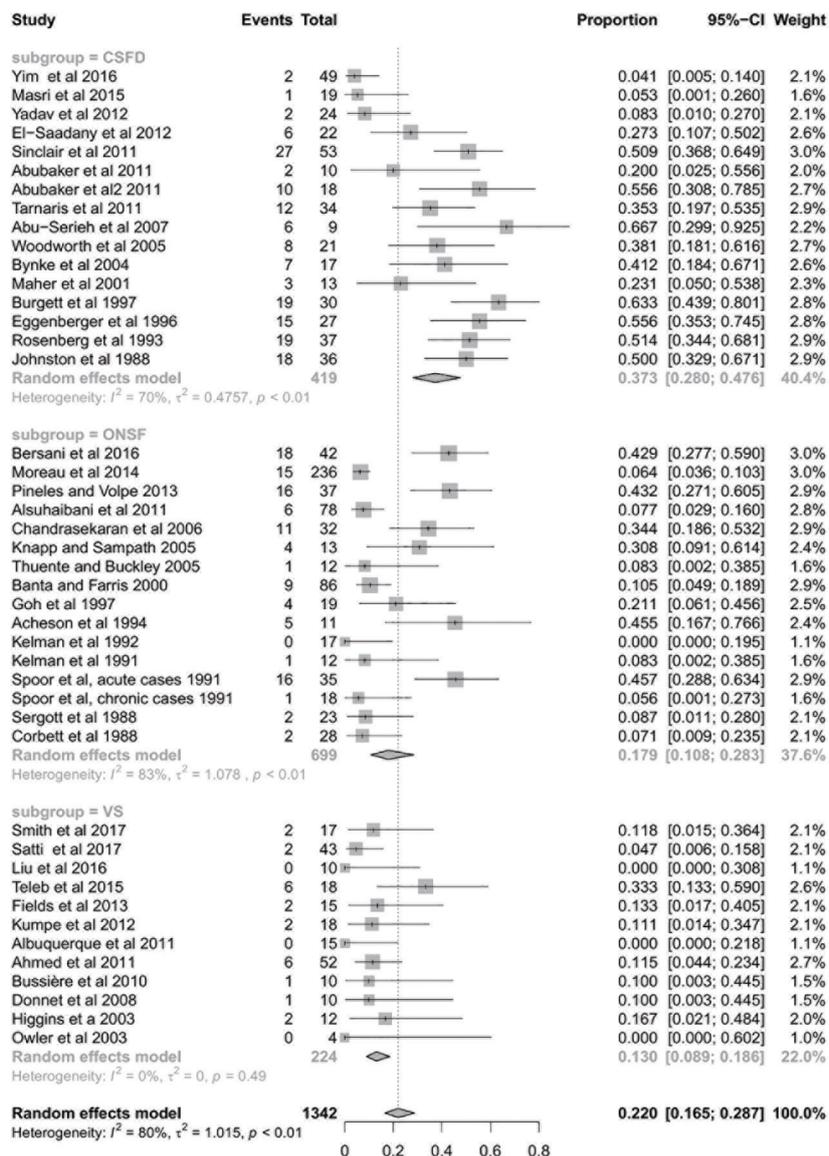


Figure 4 Forest plot of pooled subsequent procedure rate subgrouped according to CSFD, ONSF and VS. CSFD, cerebrospinal fluid diversion; ONSF, optic nerve sheath fenestration; VS, venous stenting.

CSF-diversion procedures appear to have the highest success rate for post-operative improvement in patient's experiencing headaches. Eighty-four percent of patients who underwent a CSF-diversion procedure had post-operative improvement in headaches, compared to 79% and 62% for venous sinus stenting and ONSF, respectively. However, when all treatment modalities were compared, no statistically significant difference was detected ($P=0.223$).

Venous sinus stent placement was associated with the greatest post-operative improvement in both papilloedema

and visual acuity with 86% and 69% of patients had improvements, respectively. This is in comparison to 77% and 44% of patients having post-operative improvement in papilloedema and visual acuity following ONSF, respectively. Seventy-one percent and 55% of patients had post-operative improvement in papilloedema and visual acuity following a CSF-diversion procedure, respectively. Despite the data favouring venous sinus stenting for patients with papilloedema, no statistically significant difference was detected between the treatment modalities

($P=0.192$). However, this is not the case with post-operative improvement in visual acuity, which showed a statistically significant difference favouring venous sinus stenting compared to CSF-diversion and ONSF ($P=0.037$).

Complication rates between the three treatment modalities were comparable. Optic nerve sheath fenestration had the greatest complication rate at 14%, which was comparable to the 13% complication rate for CSF-diversion procedures. Although the complication rate for venous sinus stenting was 8%, the lowest amongst the three treatment modalities, when compared against each other, no statistically significant difference was detected ($P=0.2781$).

Although the complication rate for CSF-diversion procedures is only 14%, 37% of patients undergoing such procedures require repeat or subsequent procedures. Of the 538 patients included in the meta-analysis, 157 patients underwent an additional 540 procedures, most of which were shunt revisions. With a mean follow-up time of 42 months, the high reported repeat procedure rate is concerning given that the mean age of patients undergoing a CSF-diversion procedure is 30.3 years. When compared to ONSF, which has a similar complication rate, the requirement for subsequent procedures is only 18%.

Venous sinus stent placement had a significantly lower revision rate of 13% when compared to CSF-diversion procedures and ONSF ($P<0.001$). Of the 224 patients who underwent venous sinus stenting included in the meta-analysis, only 24 patients required a subsequent procedure, of which, 12.5% had a CSF-diversion procedure. Although only 224 patients were included in the meta-analysis of dural venous sinus stent placement, the data suggests that this treatment modality is significantly more effective in improving visual acuity post-operatively and has a significantly lower requirement for additional procedures when compared to CSF-diversion procedures and ONSF.

Given the high complication rate and requirement for subsequent procedures, CSF-diversion procedures are associated with significant morbidity and cost. As such, the use of CSF-diversion procedures as the standard of treatment for medically refractive IIIH should be reconsidered. When comparing the cost of venous sinus stent placement and CSF-diversion procedure, Ahmed *et al.* found that there was no significant cost difference between the insertion of an initial venous sinus stent and initial CSF shunt. (72) Although there is no significant cost difference with respect to the initial procedure, 87% of stents placed required only one stent procedure, in comparison to only 45% of shunts requiring

only one shunt procedure (72). Given the high rate of repeat procedures, CSF-diversion procedures end up costing significantly more in the long term.

Although the results of this meta-analysis suggest the venous sinus stenting is a viable alternative to traditional CSF-diversion procedures and ONSF, there is still a limited, though growing, literature for this procedure in medically refractive IIIH. The present analysis is constrained by several limitations. These include the lack of direct comparative analyses between studies and patient matching, as baseline patient differences could be confounders in the present analysis. One would expect that patients undergoing venous stenting procedures to have venous pathology, although the extent may vary between patients and studies. The retrospective nature of the included studies means the data is susceptible to selection bias. Given the relative low number of studies of venous sinus stenting when compared to that available for CSF-diversion procedures and ONSF, combined with the retrospective nature of a meta-analysis, confirmation of these results ideally requires a randomised controlled trial before one can confidently state which treatment modality is superior in the management of medically refractive IIIH.

Conclusions

CSF-diversion procedures have traditionally been the standard of treatment for patients with medically refractive IIIH. However, the results of our meta-analysis suggest that, with its lower complication rate, lower requirement for subsequent procedures and its superiority with respect to improving visual acuity, dural venous sinus stenting may be a viable alternative to traditional surgical interventions in patients refractory to medical treatment.

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Footnote

Conflicts of Interest: All authors have completed the ICMJE uniform disclosure form (available at <http://dx.doi.org/10.21037/aes.2018.05.01>). The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related

to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

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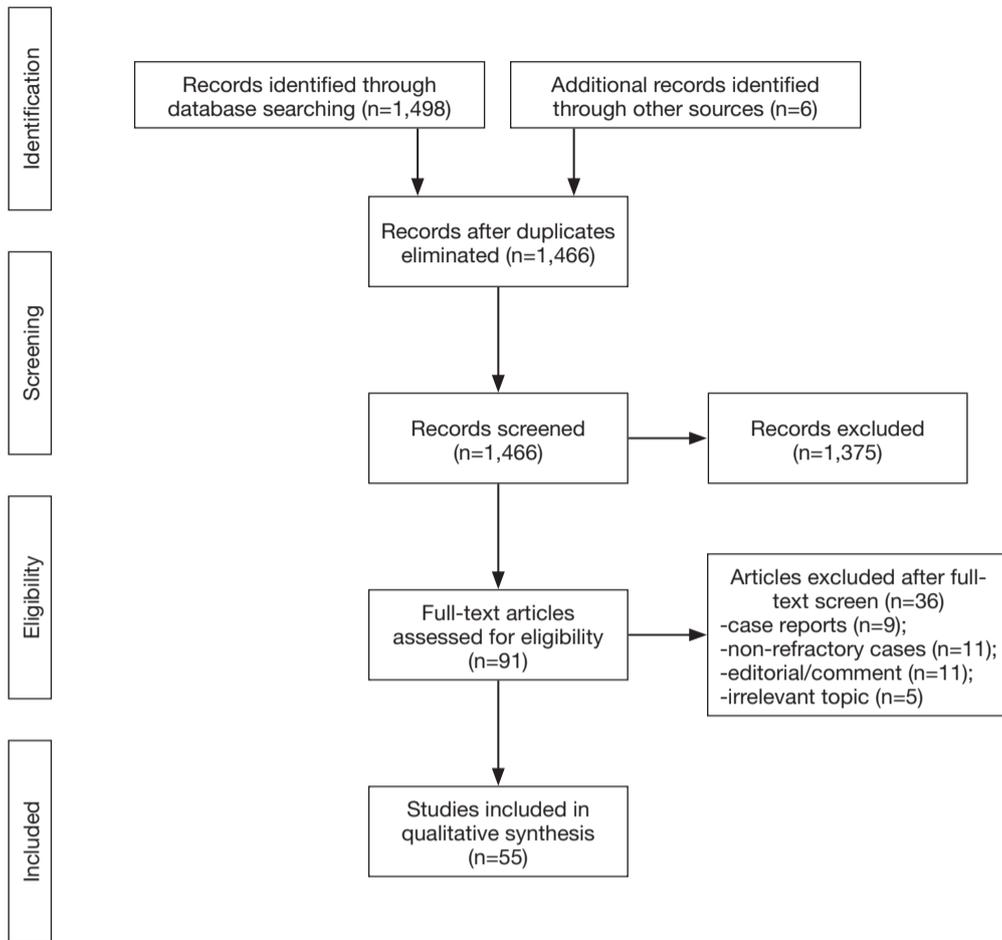


Figure S1 PRISMA flow chart demonstrating search strategy for the present systematic review and meta-analysis.

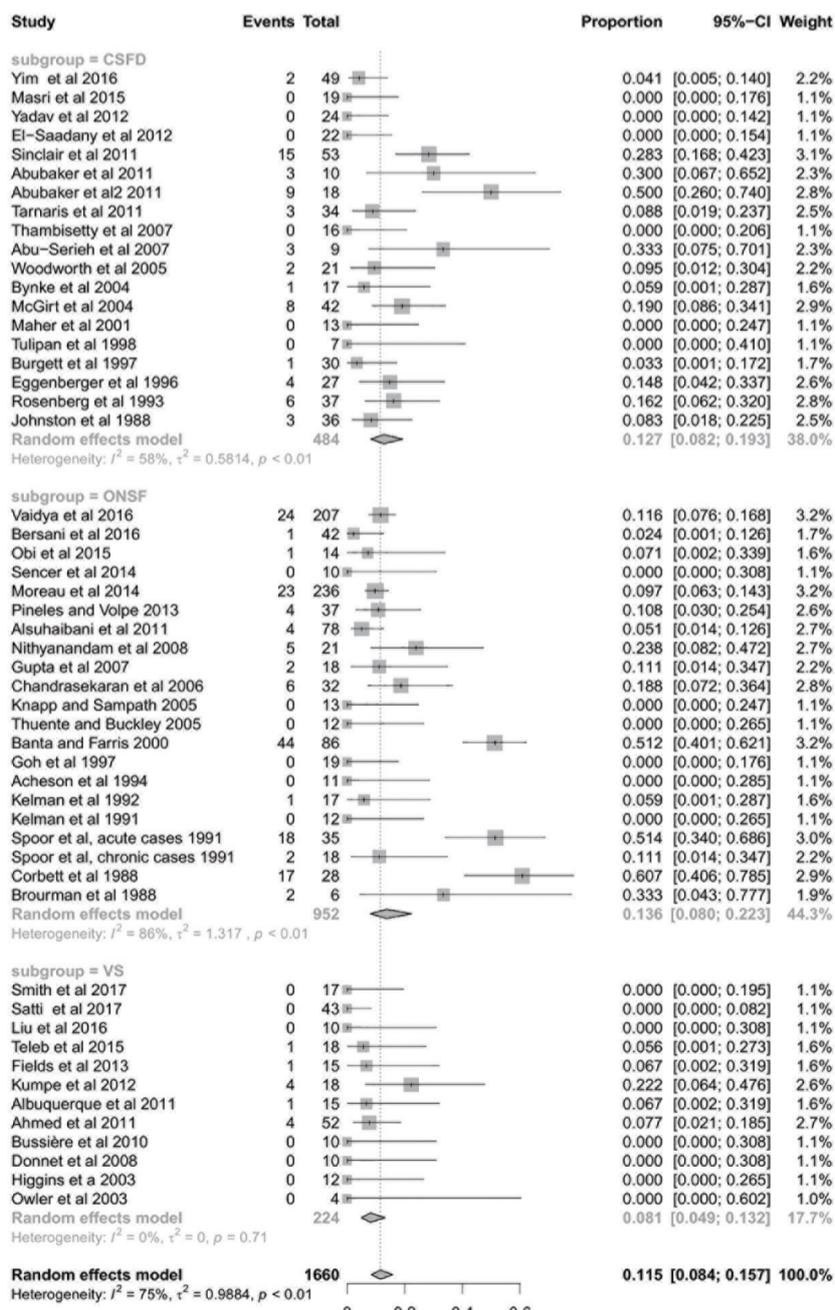


Figure S2 Forest plot of pooled complication rate subgrouped according to CSFD, ONSF and VS. CSFD, cerebrospinal fluid diversion; ONSF, optic nerve sheath fenestration; VS, venous stenting.

Table S1 CSF diversion summary

| Studies | Cases (M/F) | Age | Follow-up | BMI (kg/m ²) | CSF opening pressure (cmH ₂ O) | Primary surgery | Presenting complaint | Symptoms post-CSF diversion | | | Shunt type | Subsequent procedures | # of patients with revisions | Complications | |
|-------------------------------------|-------------|--|-------------------------------------|---|---|-----------------|-----------------------------------|-----------------------------|-------|-------|----------------|---|------------------------------|-------------------------------|--|
| | | | | | | | | HA | PAP | VAC | | | | Reasons for shunt revision | Others |
| Yim <i>et al.</i> 2016 (23) | 49 (3/46) | Mean ± SD: 33.5±12.7 years | 47 months | 37.6 | NR | NR | HA, 49; PAP, 49; VAC, NR; VFC, NR | NR/NR | NR/NR | NR/NR | VPS | NR | 2 | NR | Asymptomatic hemorrhage, 1; ventricular catheter to be in the contralateral opticocarotid cistern, 1 |
| Roth <i>et al.</i> 2015 (24) | 13 (0/13) | Mean: 27 (range, 21–31) years for BS group; Mean: 26 (range, 12–43) years for non-BS group | Mean: 86 (range, 23–181) months | BS: 43 (range, 37–47), non-BS 35 (range, 26–43) | NR | NR | HA, NR; PAP, NR; VAC, 22 | 6 remain | NR/NR | 19 | LPS, VPS | NR | NR | NR | NR |
| Rizzo <i>et al.</i> 2015 (25) | 15 (1/14) | Mean: 34 (range, 16–66) years | 2004–2011 | NR | NR | NR | HA, NR; PAP, NR; VAC, NR | NR/NR | 10/13 | NR/NR | LPS, VPS | NR | NR | NR | NR |
| Masri <i>et al.</i> 2015 (26) | 19 (11/8) | Mean: 6 years (7 months–12 years) | 2 weeks to 6 years | NR | 20 to 77 | NR | HA, 10; PAP, 12; VAC, NR | NR/NR | NR/NR | NR/NR | LPS | NR | 1 | Malfunction, 1 | None |
| Alkherayf <i>et al.</i> 2015 (27) | 7 (0/7) | Mean: 33.2 (range, 23–46) years | 5/2012–6/2013 | NR | 35.8 (range, 27 to >55) | NR | HA, NR; PAP, 5; VAC, 4; VFC, 6 | NR/NR | 4/5 | 4/4 | LP | NR | NR | NR | NR |
| Huang <i>et al.</i> 2014 (28) | 19 (1/18) | Mean ± SD: 29±13 years | 21.2 months (range, 5–1,342 days) | NR | NR | 14 | HA, 2; PAP, NR; VAC, 17; VFC, NR | NR/2 | NR/NR | 9/17 | VPS | 4 VPS revisions, 3 replacements, 1 ONSF | NR | NR | NR |
| Yadav <i>et al.</i> 2012 (29) | 24 (2/22) | Mean: 39 (range, 17–58) years | Mean: 51 (range, 18–137) months | NR | NR | NR | HA, 24; PAP, 24; VAC, 18; VFC, NR | 22/24 | NR/24 | 10/18 | LPS | 2 revisions | 2 | SF, 2 | None |
| El-Saadany <i>et al.</i> 2012 (30) | 22 (4/18) | Mean: 28.5 (range, 20–38) years | 1, 3 and 12 months | NR | NR | 22 | HA, 22; PAP, 20; VAC, NR; VFC, NR | 22/22 | 18/20 | NR/NR | LPS | 6 revisions | 6 | SO, 6 | None |
| Sinclair <i>et al.</i> 2011 (31) | 53 (3/50) | Mean ± SD: 30.3±8.5 years | Baseline, 6, 12 and 24 months | NR | 39.5 (± SD: ±8.2) | 53 | HA, 51; PAP, 44; VAC, 34; VFC, NR | 8/35 | 14/27 | 15/33 | LPS, VPS | 74 revisions | 27 | SO, 12 | CM, 2; SD, 5; others, 8 |
| Abubaker <i>et al.</i> 2011 (32) | 10 (NR) | Avail/NR (range, 25–65 years) | Mean: 48 (range, 6–96) months | NR | NR | NR | HA, 10; PAP, 7; VAC, 4; VFC, NR | 9/10 | 6/7 | 4/4 | VPS | 3 revisions | 2 | NR | CM,3 |
| Abubaker <i>et al.</i> 2011 (32) | 18 (NR) | Avail/NR (range, 25–65 years) | Mean: 48 (range, 6–96) months | NR | NR | NR | HA, 18; PAP, 17; VAC, 11; VFC, NR | 11/18 | 11/17 | 11/11 | LPS | 12 revisions | 10 | SO, 3 | SM,3; CM,6 |
| Tarnaris <i>et al.</i> 2011 (33) | 34 (2/32) | Mean: 35 (range, 27.1–42.9) years | 28.9 (±31.8) months | NR | 39.4 (range, 29.1–49.7) | 29 | HA, 34; PAP, 24; VAC, 30; VFC, NR | 20/29 | 10/24 | 12/29 | LPS, VPS | NR | 12 | SO, 1; SI, 1; LPH, 2 | AP, 1; SM, 1; CSF leak, 1 |
| Thambisetty <i>et al.</i> 2007 (34) | 16 (0/16) | Mean: 23.8 (range, 14–39) years | NR | NR | 54.1 (range, 29–60) | 16 | HA, 16; PAP, 16; VAC, 16; VFC, NR | 16/16 | NR/16 | 14/16 | LPS, VPS, ONSF | NR | None | None | None |
| Abu-Serieh <i>et al.</i> 2007 (35) | 9 (4/5) | Mean: 26.4 (range, 4–63) years | Mean: 44.3 (range, 6–110) months | NR | NR | 9 | HA, 9; PAP, 5; VAC, 2; VFC, NR | 9/9 | 3/5 | 2/2 | S-VPS | 9 revisions | 6 | SO, 1; SI, 5 | SM, 1; VD, 2 |
| Woodworth <i>et al.</i> 2005 (36) | 21 (4/17) | Mean ± SD: 42 ±10 years | 20±17 months | NR | NR | NR | HA, 21; PAP, 8; VAC, 5; VFC, NR | 21/21 | NR/8 | NR/5 | VPS | 32 revisions | 8 | SO, 21; LPH, 6 | CM, 1; CSF leak, 1 |
| Bynke <i>et al.</i> 2004 (37) | 17 (5/12) | Mean: 34 (range, 13–63) years | Mean: 78 (range, 21.6–153.6) months | 30.99 (range, 23–52.5) | 39.4 | 16 | HA, 15; PAP, 16; VAC, 8; VFC, NR | 15/15 | 16/16 | 3/8 | VPS | 9 revisions | 7 | SO, 6; SI, 2 | SM, 1 |
| McGirt <i>et al.</i> 2004 (38) | 42 (10/32) | Mean ± SD: 37±10 years | 49±31 months | NR | NR | NR | HA, 42; PAP, 25; VAC, 15; VFC, NR | 40/42 | NR/25 | NR/15 | LPS, VPS | 84 revisions | NR | SO, 55; SI, 4; LPH, 16; RP, 4 | CM, 5; TH, 3 |
| Maher <i>et al.</i> 2001 (39) | 13 (3/10) | Mean: 31.5 (range, 6–54) years | Mean: 12.4 (range, 1–38) months | NR | NR | 0 | HA, 4; PAP, NR; VAC, 11; VFC, NR | 0/4 | NR/NR | 5/11 | S-VPS | 3 revisions | 3 | SO,3 | None |
| Tulipan <i>et al.</i> 1998 (40) | 7 (NR) | NR | Mean: 9 (range, 4–17) months | NR | NR | 5 | HA, 7; PAP, 7; VAC, NR; VFC, NR | 6/7 | 5/7 | NR/NR | S-VPS | None | None | None | None |
| Burgett <i>et al.</i> 1997 (41) | 30 (2/28) | Mean: 32.9 (range, 10–68) years | Mean: 34.9 (range, 0–143) months | NR | NR | NR | HA, 17; PAP, 16; VAC, 10; VFC, NR | 14/17 | 15/16 | 6/10 | LPS | 126 revisions | 19 | SF, 13; SI, 1; LPH, 2; RP, 1 | CSF fistula, 1 |
| Eggenberger <i>et al.</i> 1996 (10) | 27 (3/24) | Mean: 28 (range, 8–51) years | Mean: 77 (range, 21–278) months | NR | NR | NR | HA, 18; PAP, 14; VAC, NR; VFC, NR | 18/18 | 9/14 | 10/NR | LPS | 66 revisions | 15 | SO, 43; SI, 1; LPH, 10; RP, 3 | AP, 1; CM, 3 |
| Rosenberg <i>et al.</i> 1993 (42) | 37 (NR) | NR | Mean: 30.9 (range, 1–180) months | NR | NR | NR | HA, 7; PAP, NR; VAC, 34; VFC, NR | NR/7 | NR/NR | 13/34 | LPS, VPS | 56 revisions | 19 | SF, 31; SI, 3; LPH, 14; RP, 2 | AP, 3; OC, 2; CSF leak, 1 |
| Johnston <i>et al.</i> 1988 (43) | 36 (10/26) | Mean: 24.7 years (6 months–54 years) | NR | NR | NR | NR | HA, 29; PAP, 34; VAC, 24; VFC, NR | NR/29 | NR/34 | NR/24 | LPS, VPS | 50 revisions | 18 | SO, 24; SI, 8; LPH, 14; RP, 1 | Others, 3 |

LPS, lumboperitoneal shunt; VPS, ventriculoperitoneal shunt; S-VPS, stereotactic ventriculoperitoneal shunt; SF, shunt failure; SO, shunt obstruction; SI, shunt infection; LPH, low-pressure headache; RP, radicular pain; AP, abdominal pain; CM, catheter migration; OC, operative complications; TH, tonsillar herniation; SM, shunt malposition; VD, valve dysfunction; SD, shunt disconnection; HA, headache; PAP, papilledema; VAC, visual acuity changes; VFC, visual field changes; M, male; F, female; NR, not reported; BS, bariatric surgery.

Table S2 Venous stenting summary

| Studies | #of cases | Sex (M/F) | Age (years) | Follow-up (months) | BMI (kg/m ²) | CSF opening pressure (cmH ₂ O) | Primary surgery | Presenting complaint | Poststenting complaints | | | | Pressure gradient (mmHg) | | Stent placement location | Complications (n) | Subsequent procedure (n) |
|-------------------------------------|-----------|-----------|----------------------------|---------------------------------|----------------------------------|---|-----------------|--|-------------------------|---|--------|-------|---|--|---------------------------------|---|--------------------------------|
| | | | | | | | | | HA | PAP | VAC | VFC | Prestent | Poststent | | | |
| Smith <i>et al.</i> 2017 (65) | 17 | 2/15 | Mean: 29.47 (range, 21–39) | Clinical: 17.5; imaging: 10.8 | Mean: 35.24 (range, 24.54–46.18) | Mean: 38.1 (range, 26–55) | NR | HA, 17; PAP, 11; VAC, 14; VFC, 13 | 16/17 | 9/11 | 10/14 | 12/13 | Mean: 23.06 | Mean: 1.18 | RTS: 17 | None | Second stenting: 2 |
| Satti <i>et al.</i> 2017 (66) | 43 | 4/39 | Mean: 34.9 | Imaging: 6.5; clinical: 13.5 | Mean: 34.8 | Mean: 35.8 | 35 | HA, 43; PAP, 28; VAC, 38; VFC, 16 | 27/39 | 13/22 | 15/35 | 6/13 | Mean: 16.74 (range, 7–46) | – | RTS: 29/42; LTS: 12/42; B/L: 10 | None | Repeat stent: 2 |
| Liu <i>et al.</i> 2016 (67) | 10 | 1/9 | Mean: 34.1 (range, 17–59) | Median: 23.4 (range, 15.7–31.6) | Mean ± SD: 41.5±9.8 | Mean: 42.5 (range, 27–55) | 0 | HA, 10; PAP, 7; VAC, 8; VFC, 10 | 9/10 | 7/7 | 5/8 | 10/10 | Mean ± SD: 39.5±14.9 | Mean ± SD: 30±13.2 | B/L: 9; unilateral stenosis: 1 | None | None |
| Teleb <i>et al.</i> 2015 (68) | 18 | 3/15 | Mean: 30 (range, 15–59) | Range, 1–45 | Mean: 36 (range, 23–59.2) | NR | 0 | HA, 18; PAP, 15; visual disturbances including: VAC, 18; VFC, NR | 5/18 | Visual disturbances including: VAC, 14/18 | 14/18 | NR/NR | Mean: 13.7 | Mean: 1.7 | RTS: 10; LTS: 3; B/L: 5 | Deep vein thrombosis: 1 | Re-stenosis and retreatment: 6 |
| Fields <i>et al.</i> 2013 (11) | 15 | 0/15 | Mean: 34 (range, 20–56) | Mean: 14 (range, 1–49) | Mean: 39 (range, 30–73) | NR | 9 | HA, 15; PAP, 15; VAC, 14; VFC, NR | 10/15 | 15/15 | 13/13 | NR/NR | Mean: 24 (range, 13–40) | Mean: 4 (range, 0–9) | RTS: 8; LTS: 4; B/L: 3 | Femoral pseudoaneurysm: 1 | CSF diversions [VPS]: 2 |
| Kumpe <i>et al.</i> 2012 (16) | 18 | 6/12 | Mean: 37.9 (range, 16–62) | Mean: 43.7 (range, 11–136) | Mean: 31.6 (range, 22.6–38) | Mean: 37.9 (range, 25–55); NR in 6 | 8 | HA, 12; PAP, 16; VAC, 17; VFC, NR | 10/12 | 15/16 | NR/17 | NR/NR | Mean: 21.1 (range, 10.5–39) | Mean: 2.5 (range, 0–7) | RTS, 12; LTS, 7 | Subdural hematoma: 2; UTI: 1; syncope: 1 | Repeat stent: 2 |
| Albuquerque <i>et al.</i> 2011 (18) | 15 [18] | 3/12 | Mean: 31 (range, 12–51) | Mean: 20 (range, 2–40) | NR | NR | NR | HA, 15; PAP, NR; VAC, NR; VFC, NR | 12/15 | NR/NR | NR/ NR | NR/NR | NR | NR | RTS, 9; LTS, 6 | Retroperitoneal hematoma: 1 | None |
| Ahmed <i>et al.</i> 2011 (17) | 52 | 5/47 | Mean: 34 (range, 10–64) | Mean: 24 (range, 2–108) | >30 in 47 patients | Mean: 32.2 (range, 25–73); NR in 9 | 43 | HA, 43; PAP, 46; VAC, 13; VFC, 30 | 40/43 | 46/46 | 9/13 | 23/30 | Mean: 19.1 (range, 4–41) | Mean: 0.6 (range, 0–14) | RTS, 36; LTS, 16; NR, 4 | Subdural hematoma: 2; transient hearing loss: 2 | Repeat stent: 6 |
| Bussière <i>et al.</i> 2010 (20) | 10 [13] | 0/10 | Range, 16–65 | Mean: 20.1 (range, 4–60) | Mean: 35.9 (range, 27.2–47.4) | Range, 25–50 | 10 | HA, 10; PAP, 9; VAC, 8; VFC, 4 | 10/10 | 9/9 | 7/8 | NR/4 | Mean: 28.3 (range, 11–50); >10 in 2 cases | Mean: 11.25 (range, 2–23); NR in 2 cases | RTS, 8; LTS, 2 | None | CSF diversions (VPS): 1 |
| Donnet <i>et al.</i> 2008 (9) | 10 | 2/8 | Mean: 41.8 (range, 28–60) | Mean: 17 (range, 6–36) | Mean: 27.3 (range, 22–37) | Mean: 40.2 (range, 29–59) | 10 | HA, 10; PAP, 10; VAC, 10; VFC, NR | 8/10 | 10/10 | NR/10 | NR/NR | Mean: 19.1 (range, 12–34) | NR | RTS, 7; LTS, 2; B/L, 1 | None | Contralateral stents: 1 |
| Higgins <i>et al.</i> 2003 (19) | 12 | 0/12 | Mean: 33 (range, 19–52) | Mean: 14.1 (range, 2–26) | Mean: 36.9 (range, 29–45) | Mean: 33.7 (range, 25–46) | 7 | HA, 12; PAP, 8; VAC, 12; VFC, NR | 7/12 | 5/8 | 7/12 | NR/NR | Mean: 18.9 (range, 8–37) | Mean: 5.75 (range, 2–15) | NR | None | Contralateral stents: 2 |
| Owler <i>et al.</i> 2003 (73) | 4 [9] | 1/3 | Mean: 27.3 (range, 17–38) | Mean: 9.25 (range, 5–12) | Mean: 30 (range, 23–38) | Mean: 28.7 (range, 22–35); NR in 1 | 1 | HA, 4; PAP, 4; VAC, 4; VFC, 3 | 4/4 | 4/4 | 4/4 | 3/3 | Mean: 18.8 (range, 12–25) | Mean: 0.25 (range, 0–1) | RTS, 3; LTS, 1 | None | None |

M, indicates male; F, female; HA, headache; PAP, papilledema; VAC, visual acuity changes; VFC, visual field changes; RTS, right transverse sinus; LTS, left transverse sinus; B/L, bilateral; UTI, urinary tract infection; VPS, ventriculoperitoneal shunt; NR, not reported.

Table S3 ONSF summary

| Studies | Cases (M/F) | #of eyes | Age, mean [range] (years) | Follow-up | BMI, mean (kg/m ²) | CSF opening pressure (cmH ₂ O) | Surgery (n) | | | Presenting complaint | Post-ONSF complaints | | | | | | Complications (n) | Subsequent procedures (n) | |
|--|-------------|-----------------|---------------------------|--|--------------------------------|---|-------------|------------|---------|-------------------------------------|----------------------|---|----------|-----------------|---------|---|---|---|----|
| | | | | | | | Bilateral | Unilateral | Primary | | HA | PAP | VAC | | VFC | | | | |
| | | | | | | | | | | | | | Improved | Improved/stable | | | | | |
| Vaidya <i>et al.</i> , 2016 (44) | 104 (4/100) | 207 | Mean: 28.8 | 6 months | NR | Mean: 39.85 | 103 | 1 | 0 | HA, NR; PAP, 173; VAC, NR; VFC, N/A | NR/NR | At 1 week: 102/173; at 1 month: 90/173; at 6 months: 94/173 | | NR/NR | NR/NR | At 1 week: 148/148; at 1 month: 128/128; at 6 months: 128/128 | | Transient diplopia: 7; efferent pupillary dysfunction: 17 | NR |
| Bersani <i>et al.</i> , 2016 (45) | 42 (10/32) | 64 | Mean: 26.1 | 6–12 weeks | 42.55 (18 subjects) | NR | NR | NR | NR | HA, NR; PAP, NR; VAC, 64, VFC, 29 | NR/NR | NR/NR | 64/64 | – | 28/29 | A left exotropia: 1 | Repeat ONSD of the operated eye: 3 (7%); ONSD of the contralateral eye: 12 (29%); or both: 3 (7%) | | |
| Obi <i>et al.</i> , 2015 (46) | 14 (5/9) | 31 | 35.5 [16.5–61] | Mean: 26 months (range, 2 months–6 years) | 10 had a BMI >30 | NR | 11 | 3 | 0 | HA,13; PAP,14; VFC, N/A | 7/13 | 14/14 | 7/29 | 18/29 | 14/29 | Transient diplopia: 2; transient ocular discomfort: 1 | NR | | |
| Sencer <i>et al.</i> , 2014 (47) | 10 (1/9) | 10 | 34.1 [9–49] | Mean: 28.4 (range, 8–55) months | NR | NR | 0 | 10 | 9 | HA, 7; PAP, 9; VAC, 9; VFC, 9 | 4/7 | 7/9 | 7/9 | 8/9 | 8/9 | None | NR | | |
| Moreau <i>et al.</i> , 2014 (48) | 236 (NR) | 455 | NR | Mean: 18.7 months (range, 1 week–10 years) | NR | NR | NR | NR | NR | HA, NR; PAP, NR; VAC, 448; VFC, 227 | NR/NR | NR/NR | 75/448 | 429/448 | 142/227 | Diplopia: 15; dellen: 2; esotropia: 4; exotropia: 2 | Repeat ONSD: 15 | | |
| Pineles and Volpe, 2013 (49) | 37 (5/32) | 50 | 33 [19–74] | Mean: 48.2 (range, 1–160) months | NR | NR | 13 | 24 | – | HA, NR; PAP, NR; VAC, 37; VFC, 16 | NR/NR | NR/NR | 8/37 | 28/37 | 16-6 | Tonic pupil: 2; conjunctival abscess: 1; diplopia: 1 | CSF diversion: 8; repeat ONSD: 8 | | |
| Alsuhaibani <i>et al.</i> , 2011 (50) | 78 (12/66) | 88 | 32 [13–57] | 2 weeks–12 months | NR | NR | 10 | 68 | 78 | HA, NR; PAP, NR; VAC, NR; VFC, NR | NR/NR | NR/NR | NR/NR | NR/NR | NR/NR | Transient diplopia: 3; large cyst formation at the site of the surgery: 1 | CSF diversion: 6 | | |
| Nithyanandam <i>et al.</i> , 2008 (51) | 21 (6/15) | 41 ^a | 29.5 [18–48] | 3–29 months | NR | NR | 19 | 2 | 21 | HA, NR; PAP, NR; VAC, 34; VFC, 34 | NR/NR | NR/NR | 17/34 | 32/34 | 22/34 | Transient pupillary atony: 3; transient diplopia: 1; orbital cellulitis: 1 | NR | | |
| Gupta <i>et al.</i> , 2007 (52) | 18 (4/14) | NR | 35.8 [30–42] | 3–45 months | 24.4 | Mean: 26.8 (range, 25–70) | 0 | 18 | 15 | HA, 18; PAP, NR; VAC, 36; VFC, NR | NR/18 | NR/NR | 30/36 | 36/36 | NR/NR | Synechiae: 2 | NR | | |
| Chandrasekaran <i>et al.</i> , 2006 (53) | 32 (3/29) | 51 | 33.4 [17–65] | 27.6 (range, 0–120) months | NR | NR | 18 | 14 | 25 | HA, NR; PAP, 32; VAC, 31; VFC, 39 | NR/NR | NR/32 | 13/31 | 30/31 | 13/39 | Transient diplopia: 3; anisocoria: 2; disc hemorrhage: 1 | CSF diversion: 11 | | |
| Knapp and Sampath, 2005 (54) | 13 (4/9) | 23 | 26.5 [14–49] | 9.6 (range, 1–32) months | NR | NR | 10 | 3 | 11 | HA, NR; PAP, NR; VAC, 27; VFC, 24 | NR/NR | NR/NR | 4/27 | 27/27 | 18/24 | None | Repeat ONSD: 4 | | |
| Thuente and Buckley, 2005 (55) | 12 (6/6) | 17 | 10.1 [4.4–16] | Mean: 39.6 (range, 2.4–105.3) months | NR | Mean: 33 (range, 13–47) | 5 | 7 | 12 | HA, 8; PAP, 12; VAC, 17; VFC, 17 | 5/8 | 12/12 | 7/17 | 17/17 | 6/17 | None | Repeat ONSD: 5; CSF diversion: 1 | | |
| Banta and Farris, 2000 (56) | 86 (13/73) | 158 | 32.1 | Mean: 20 (range, 1–108) months | NR | NR | 72 | 14 | 86 | HA, 61; PAP, 86; VAC, 158; VFC, 81 | 8/61 | NR/86 | NR/158 | 148/158 | 71/81 | Diplopia: 30; dellen: 6; anisocoria:6; orbital apex syndrome: 1; presumed traumatic optic neuropathy: 1 | Repeat ONSD + CSF diversion: 4; CSF diversion: 2; repeat ONSD: 3 | | |
| Goh <i>et al.</i> , 1997 (57) | 19 (6/13) | 29 | 33.1 [16–52] | Mean: 15.7 (range, 1–50) months | NR | NR | 10 | 9 | 19 | HA, NR; PAP, NR; VAC, 29; VFC, 21 | NR/NR | NR/NR | 4/29 | 26/29 | 10/21 | None | Repeat ONSD: 4 | | |
| Acheson <i>et al.</i> , 1994 (58) | 11 (4/7) | 15 | 37.1 [23–53] | Mean: 24 (range, 12–84) months | NR | NR | 4 | 7 | 7 | HA, NR; PAP, NR; VAC, 15; VFC, 15 | NR/NR | NR/NR | 8/15 | 13/15 | 8/15 | None | CSF diversion: 4; subtemporal decompression: 1 | | |
| Kelman <i>et al.</i> , 1992 (59) | 17 (2/15) | 21 | 40 [16–73] | Mean: 17 (range, 11–26) months | NR | NR | 4 | 13 | 13 | HA, 10; PAP, 17; VAC, 21; VFC, 21 | 9/10 | 17/17 | 14/21 | 20/21 | 20/21 | Orbital hematoma: 1 | None | | |
| Kelman <i>et al.</i> , 1991 (60) | 12 (1/11) | 15 | 38 [31–48] | Mean: 31 (range, 3–72) months | NR | NR | 3 | 9 | 0 | HA, 7; PAP, 12; VAC, 24; VFC, 24 | 6/7 | NR/12 | 15/24 | 24/24 | 21/24 | None | Repeat ONSD: 1 | | |
| Spoor <i>et al.</i> , 1991 (61) | | | | | | | | | | | | | | | | | | | |
| Acute cases | 35 (4/31) | 69 | 32.3 [6–72] | Mean: 18.1 (range, 2–48) months | NR | NR | 21 | 14 | 33 | HA, 7; PAP, 35; VAC, 69; VFC, 69 | NR/7 | NR/35 | 29/69 | 69/69 | 68/69 | Primary surgery: transient diplopia, pupillary dysfunction, hypotonia, peripapillary hemorrhages: 2; repeat surgery: pupillary dysfunction: 2; peripapillary hemorrhages: 1; dellen, chorioretinal scarring, late failure: 13 | Repeat ONSD: 16 | | |
| Chronic cases | 18 (5/13) | 32 | 32.7 [7–57] | Mean: 14.6 (range, 3–46) months | NR | NR | 13 | 5 | 13 | HA, 3; PAP, 18; VAC, 32; VFC, 24 | NR/3 | NR/18 | 9/32 | 32/32 | 7/24 | | Repeat ONSD: 1 | | |
| Sergott <i>et al.</i> , 1988 (62) | 23 (1/22) | 29 | 38.1 [18–63] | Mean: 21.5 (range, 3–45) months | NR | NR | 6 | 17 | 17 | HA, 17; PAP, 23; VAC, 29; VFC, 29 | 13/17 | 12/23 | 21/29 | 28/29 | 29/29 | Perilimbal conjunctival bleb: 1; horizontal diplopia: 1 | Repeat ONSD: 2 | | |
| Corbett <i>et al.</i> , 1988 (63) | 28 (8/20) | 40 | 29.3 [14–62] | Mean: 26.9 (range, 0–90) months | NR | NR | 12 | 16 | 26 | HA, 17; PAP, 28; VAC, 40; VFC, 38 | 11/17 | 24/28 | 12/40 | 34/40 | 21/38 | Tonic pupil: 16; retrobulbar hemorrhage: 1 | CSF diversion: 2 | | |
| Brouman <i>et al.</i> , 1988 (64) | 6 (0/6) | 10 | 38.5 [28–62] | 4–11 months | NR | NR | 4 | 2 | 4 | HA, NR; PAP, 6; VAC, 10; VFC, NR | NR/NR | 4/6 | 3/10 | 10/10 | NR/NR | Transient diplopia: 1; transient atonic pupil: 1 | NR | | |

^a, seven eyes were excluded from analysis due to no light perception. HA, indicates headache; PAP, papilledema; VAC, visual acuity changes; VFC, visual field changes; M, male; F, female; NR, not reported; N/A, not available; ONSD, optic nerve sheath decompression.