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**A meta-analysis of the effects of Venous super-drainage in Deep Inferior Epigastric
Perforator flaps for breast reconstruction.**

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A meta-analysis of the effects of Venous super-drainage in Deep Inferior Epigastric artery Perforator flaps for breast reconstruction.

Abstract

Introduction

Venous congestion is the most common vascular complication of the Deep Inferior Epigastric artery Perforator (DIEP) flaps. Adding a second venous drainage by anastomosing a flap vein and a recipient vein (super-drainage) is considered the solution of choice. Evidence to support this procedure, had not yet been confirmed by an analysis of the literature. We aimed to provide this evidence.

Materials and Methods

We searched the literature (MedLine, Scopus, EMBASE, Cochrane Library, and Google Scholar), for studies discussing venous congestion and venous super-drainage in DIEP flap for breast reconstruction. 13/35 articles compared results between one or two venous anastomoses. Meta-analysis was performed following PRISMA guidelines . Pooled risk ratio (RRs) for congestion, fat necrosis, partial necrosis, and total necrosis with corresponding 95% confidence intervals (CI) were calculated using a fixed-effect model with the Mantel–Haenszel method. The need to return to surgery (95% CI) was estimated with a random effect model using the DerSimonian & Liard method.

Results

We showed a statistically significant advantage of super-drainage to reduce the venous congestion of the flap (RR: 0.12, 95% CI: 0.04 – 0.34, p -value < 0,001), partial flap necrosis (RR: 0.50, 95% CI: 0.30 - 0.84, p -value 0,008), total flap necrosis (RR: 0.31, 95% CI: 0.11 - 0.85, p -value 0.023), and the need to take the patient back to surgery for perfusion-related complications (RR: 0,45, 95% CI: 0.21 - 0.99, p value 0.048).

Conclusions

Performing a second venous anastomosis between the SIEV and a recipient vein (venous superdrainage) reduces venous congestion and related complications in DIEP flaps for breast reconstruction.

Keywords: DIEP, venous congestion, meta-analysis, super-charging, super-drainage

INTRODUCTION

The Deep Inferior Epigastric artery Perforator (DIEP) flap, is considered the gold standard for autologous breast reconstruction.¹ Flap failure is reported to occur in 1 to 5 % of cases.² However, the frequency of partial failures and their consequences is higher.³ The unsatisfactory results are frequently due to perfusion-related complications, developing in up to 17% of DIEP flaps.⁴ Venous congestion is the most common of the above-mentioned perfusion-related complications, with a reported incidence ranging from 2% to 20%^{5,6} representing one of the main causes of fat necrosis, partial flap necrosis and, less frequently, complete necrosis of the flap.³ The etiology of venous congestion is multifactorial⁷⁻¹² with reversible and non-reversible causes.

When the causes of venous congestion are not reversible intraoperatively, treatment of the congestion itself is necessary and usually consists in increasing venous drainage of the flap by adding a second venous anastomosis between the flap and a recipient vein. The term flap supercharge or, semantically more correctly, venous super-drainage is usually employed.

Several papers have been published highlighting the problem of DIEP's venous congestion, and describing the surgical solutions available. **Although there is no controversy regarding the use of flap super-drainage, which is the standard solution in case of venous congestion, evidence of the efficacy of this procedure on flap necrosis and vascular complications had not yet been reported.**

To try to clarify the matter, we conducted a literature search statistically evaluating with a meta-analysis the relevant studies looking for evidence on the role of DIEP flap's venous super-drainage in decreasing flap complications.

MATERIALS AND METHODS

We performed a meta-analysis of the literature evaluating the role of a second venous anastomosis to prevent the complications of DIEP flap's congestion. A literature search was conducted using MedLine, Scopus, EMBASE, Cochrane Library, and Google Scholar to identify and include all citations from 1994 to June 2019. To reduce inclusion bias, two of the Authors (VP and FG) performed, separately, the initial article search and selection with pertinent keywords ("Deep inferior epigastric perforator" OR "DIEP" AND "flap" [All Fields] AND venous congestion [All Fields] AND ("additional anastomosis"[Subheading] AND "superficial inferior epigastric vein"[All Fields] OR "SIEV"[All Fields] AND "supercharging" OR "superdrain" OR "super-drainage") (Figure 1)

Duplicated articles, isolated abstracts, case reports, correspondence and letters were excluded and only full-text articles in English regarding DIEP flaps and venous congestion were considered.

We followed the guidelines of the Preferred Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA statement)¹³.

We considered eligible the retrieved publications based on PICOS criteria (Patients, Interventions, Control/comparison, Outcomes, and Study Design), and analyzed by meta-analysis the incidence of perfusion-related complications: venous congestion, flap necrosis (total and partial), fat necrosis and need to take the patient back to surgery for perfusion-related complications. The most relevant data collected from the publications found were: first author, year of publication, number of patients and events, type of the second anastomosis and recipient vein, and operative time.

Statistical Analysis

The Mantel-Haenszel test in R was used to calculate Pooled RRs and corresponding 95% confidence intervals (CI) for perfusion-related complications. The heterogeneity among studies

was analyzed by using the I^2 test. We used a fixed-effects model for I^2 ranging from 0 to 30%; in the case of $I^2 > 30\%$, we applied a random effect model using the DerSimonian & Liard method.

For each test p value $\leq 0,05$ was set for significance

RESULTS

After exclusion of non-relevant papers, we identified 35 eligible articles and performed a qualitative evaluation following the guidelines of the Preferred Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA statement).¹³ Of these articles, 13 compared results between one or two venous anastomoses in DIEP flaps for breast reconstruction^{3,14-25} and were included in our study. All the studies that we reviewed were retrospective, except the one by Ayestaray,¹⁶ which is prospective and randomized.

Overall, our study analyzed 3094 DIEPs, of which, 1279 received a second venous anastomosis ("case group"); and 1815 received only one venous anastomosis ("control group"). Venous congestion occurred in 38 out of 1121 flaps, with an overall mean rate of 3.4%. The venous congestion rate in each study ranged from 0.9% to 36.5%. Regarding other perfusion-related complications, our literature review revealed 1.5 % total flap loss (34/2278) (ranging from 0% to 9.6%), 6% partial flap loss (78/1294) (0% to 28.8%), 11.9% fat necrosis (217/1824) (7.2% to 34.6%), 11.1% take back to surgery for perfusion-related complications (187/1678) (0.9% to 21.2%).

Of the 13 articles^{3,14-25} included in our meta-analysis, 12^{3,14-18,20-25} reported the second anastomosis to have been performed between the SIEV and a recipient vein of the thorax, while in one study¹⁹ the second anastomosis was done using as flap vein the SIEV or a second DIEV, without additional information being provided. In all the articles included in our systematic review, performing an adjunctive anastomosis variably increased the operation time from no difference to 105 minutes (TABLE 1).

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3 In more detail we found that anastomosing the SIEV of the flap with a recipient vein provided a
4 statistically significant advantage in terms of venous congestion (RR: 0.12, 95% CI: 0.04 – 0.34) *p*-
5 value < 0,001) (Figure 2) and to prevent flap necrosis both partial (RR: 0.50, 95% CI: 0.30 - 0.84, *p*-
6 value 0,008) (Figure 3) and total (RR: 0.31, 95% CI: 0.11 - 0.85, *p*-value 0.023) (Figure 4). A second
7 venous drainage showed a statistically significant advantage also regarding the need to take the
8 patient back to surgery for perfusion-related complications (RR: 0,45, 95% CI: 0.21 - 0.99, *p* value
9 0.048) (Figure 5). Conversely, the advantage of a second venous anastomosis was not significantly
10 beneficial to prevent fat necrosis RR: 0.83, 95% CI: 0.63 - 1.10 reporting a *p*-value 0.19). (Figure 6)
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25 **DISCUSSION**

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30 Although the DIEP flap is considered the gold standard for breast reconstruction, venous
31 congestion can cause complete or partial flap failure and several other complications.

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33 Our work, by reviewing the updated pertinent literature, aimed to provide evidence on
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35 the effect of super-drainage in terms of venous congestion and related complications. In case of
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37 intraoperative venous congestion, the reversible causes can be identified and corrected. In
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39 particular, anastomotic failure requires resection of the anastomotic site, removal of any
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41 thrombus present, rinsing of the donor and recipient vessels with a heparinized solution (and,
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43 depending on the clinical situation, time since the beginning of occlusion and kind of flap, flap
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45 perfusion with thrombolytic agents), and vessels' re-anastomosis. Vessel kinking can be solved by
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47 repositioning the entire pedicle of the flap by supporting it with a piece of fat or muscle, by
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49 stabilizing it with fibrin glue or by applying a stitch between the kinked vessel and the surrounding
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51 tissue to straighten its course. A mild to moderate torsion of a long pedicle can be solved simply
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53 by repositioning of the pedicle and therefore redistributing the torsion on its entire length. In
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more severe cases the anastomosis will need to be redone. If, in a large flap, the venous congestion is stable and only involves the distal portion of the flap, because of insufficient drainage despite effective outflow through the pedicle, resection of that area may be sufficient.

The second venous anastomosis, aimed to obtain a super-drainage of the flap, can be performed

a) between the deep venous system of the flap and a comitant vein of the recipient vessels (comitant DIEV-IMV)^{19,26}, b) between the deep venous system of the flap and a vein of the thorax (comitant DIEV-vein of thorax)²⁷, c) between the SIEV and the DIEV or to one of its side-branches (SIEV-DIEV)^{1,5,17,28,29}

d) between SIEV and a vein of the thorax^{3,14-22,24}.

In physiologic conditions, the dominant venous drainage to the lower abdominal skin and fat is provided by the superficial venous system through the inferior epigastric vein (SIEV).^{5, 15} All the papers included in our meta-analysis used the SIEV as a second vein of the flap to increase venous output. Table 1. The SIEV can be anastomosed, as said, to a vein of the deep venous system of the flap (intra-flap anastomosis) 1,5 or a vein of the patient's chest (extra-flap anastomosis).¹⁴

Unfavorable anatomy of the venous system is considered among the most important, not-reversible, causes of venous congestion. Several studies^{5,6,10,30} attribute the clinical manifestations of venous congestion and the so-called perfusion-related complications to the lack of sufficient venous drainage from all areas of the flap (superficial, deep, ipsilateral, contralateral). Not all the factors related to the flap's venous anatomy can be modified by the surgeon. Therefore accurate planning based on vascular anatomy is of utmost importance to prevent venous congestion of the flap. Computed tomographic angiography and magnetic resonance angiography permit to visualize the arterial system and, even more importantly, the venous connections between the deep and superficial systems, allowing to choose the most favorable perforators for optimal perfusion and

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3 drainage.^{7,10} The ideal perforator will have a large caliber, a central position in the flap and
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5 numerous connections between the deep and superficial venous systems.³¹
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8 Once venous congestion occurs, and intraoperatively the reversible causes have been excluded, it
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10 is necessary to treat the venous congestion itself, by venous super-drainage.
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13 One previously published meta-analysis, from Lee et al³² showed that the use of super-drainage
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15 has a statistical advantage over venous congestion but only a trend toward a decreased risk of
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17 congestion-related complications. Further studies were, therefore, suggested. A detailed
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19 comparison with that previous meta-analysis³², is presented in TABLE 2. Our analysis includes the
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21 six articles examined by Lee and seven additional ones, reporting on large groups of patients^{22,23}
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23 and on a prospective randomized study¹⁶. (TABLE 1). Statistical methods were comparable but the
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25 larger amount of data, allowed us to achieve statistical evidence that the use of super-drainage,
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27 employing a second venous anastomosis between the SIEV and a recipient vein, reduces venous
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29 congestion, prevents partial and total flap necrosis, and the need to take the patient back to
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31 surgery. Unfortunately, the available data did not show a statistically significant advantage of a
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33 second venous anastomosis to prevent fat necrosis, a common complication (6% to 17.4%) in DIEP
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35 flaps.
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42 However, from the results of our meta-analysis, there seems to be an advantage also in
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44 performing a second venous anastomosis in flaps without venous congestion. Therefore, the
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46 choice of performing routinely a second venous anastomosis should be considered.
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52 A comparison between the super-drainage technique consisting in anastomosis of the SIEV to a
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54 recipient vein (the only one for which we have demonstrated a statistical advantage), and other
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56 techniques, such as DIEV to recipient vein, intermittent SIEV catheter drainage, or the application
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of leeches, was not feasible, based on the data available in the literature. The same problem exists regarding which vein the SIEV was anastomosed (e.g., retrograde IMV vs cephalic turndown).

A limitation of our study consists in the difficulty to achieve solid conclusions when working on mainly retrospective articles , none of which with an evidence level I or II (TABLE 1). However, our study follows the PRISMA guidelines ¹³, which warrant the quality of a meta-analysis.

Our study is the first to report statistically significant results on the subject of super-drainage. Our meta-analysis of the literature demonstrated the usefulness of venous superdrainage when performing DIEP flaps in postmastectomy breast reconstruction, an elective surgery burdened with relevant emotional implications for the patient. The use of an adjunctive venous anastomosis as a preventive measure could be considered.

CONCLUSIONS

In conclusion, our meta-analysis shows that venous super-drainage, i.e. performing a second venous anastomosis between the superficial venous system and a recipient vein, provides a statistically significant advantage in terms of venous congestion and related complications in DIEP flaps for breast reconstruction.

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CRediT authorship contribution statement

Marco Pignatti: Conceptualization, Data curation, Formal analysis, Writing - original draft, Writing - review & editing. Valentina Pinto: Conceptualization, Data curation, Formal analysis, Writing - original draft.

Federico A. Giorgini: Conceptualization, Data curation.

Maria Elisa Lozano Miralles: Conceptualization, Data curation, Formal analysis. Salvatore D'Arpa: Conceptualization, Data curation Writing - original draft, review and editing.

Riccardo Cipriani: Conceptualization, Formal analysis, Writing - review & editing.

Giorgio De Santis: Conceptualization, Formal analysis, Writing - original draft, Writing - review & editing.

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LEGENDS to Figures and Tables

Figure 1

Flow diagram that shows selection of articles.

Figure 2

Venous congestion in DIEP flaps.

Articles and related forest plots dealing with the complication

VC: number of flaps presenting Venous Congestion

The column "Cases" represents the total number of flaps where an adjunctive second vein anastomosis (SIEV-recipient vein) was performed. The column "Controls" represents the total number of flaps where a single vein anastomosis (DIEV-main vein) was performed

Figure 3

Partial necrosis in DIEP flaps.

Articles and related forest plots dealing with the complication

PN: number of flaps presenting Partial Necrosis

Figure 4

Total necrosis in DIEP flaps.

Articles and related forest plots dealing with the complication

TN: number of flaps presenting Total Necrosis

Figure 5

Need to take back to surgery for perfusion-related complications in DIEP flaps.

Articles and related forest plots dealing with each complication

TB: number of flaps needing Take Back to surgery

Figure 6

Fat necrosis

Articles and related forest plots dealing with the complication

FN: number of flaps presenting Fat Necrosis

TABLE 1.

Studies included in our meta-analysis

OT: operating time, NA: not available, TAV: thoracoacromial vein, TDV: thoracodorsal vein, ICV: intercostal vein, IMV: internal mammary vein, LTV: lateral thoracic vein, IMVr: internal mammary vein retrograde branch, IMVp: internal mammary vein perforator, SCV: scapular circumflex vein, SUV: subscapular vein, AV: axillary vein

“Cases”: flaps with a second venous anastomosis. “Controls”: flaps with a single venous anastomosis

In bold are the 7 studies that were not analyzed in the previously published meta-analysis by Lee³²,

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TABLE 2.

Comparison between our meta-analysis and the one from Lee and Mun

RR: Risk Ratio. Cases: flaps with a second venous anastomosis. Controls: only one venous anastomosis

In bold are the 7 studies that were not analyzed in the previously published meta-analysis by Lee³², “Cases”: flaps with a second venous anastomosis. “Controls”: flaps with a single venous anastomosis

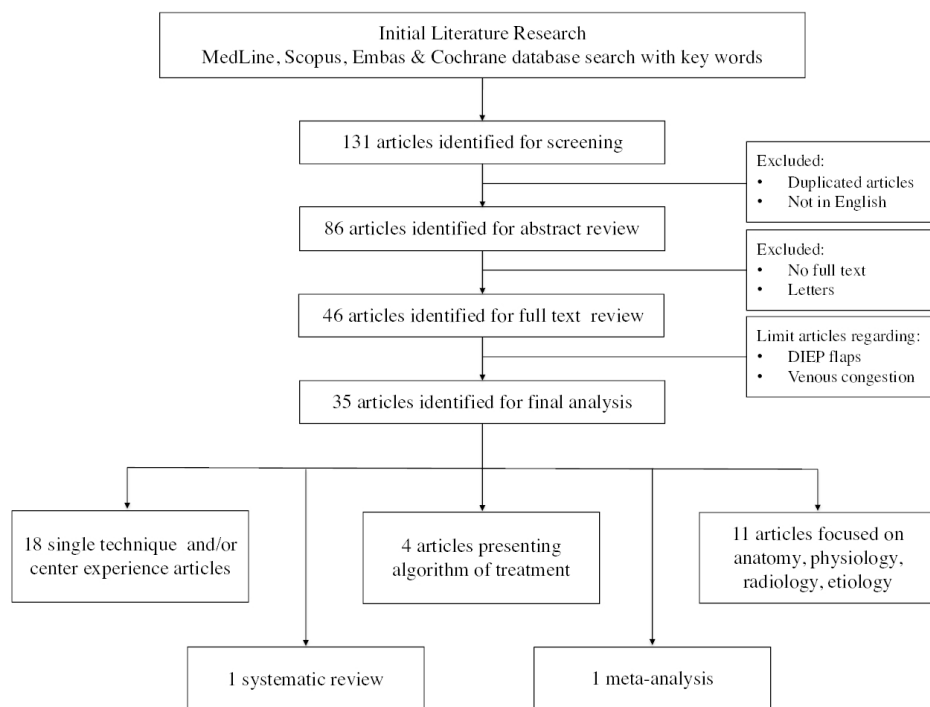


Figure 1

355x266mm (96 x 96 DPI)

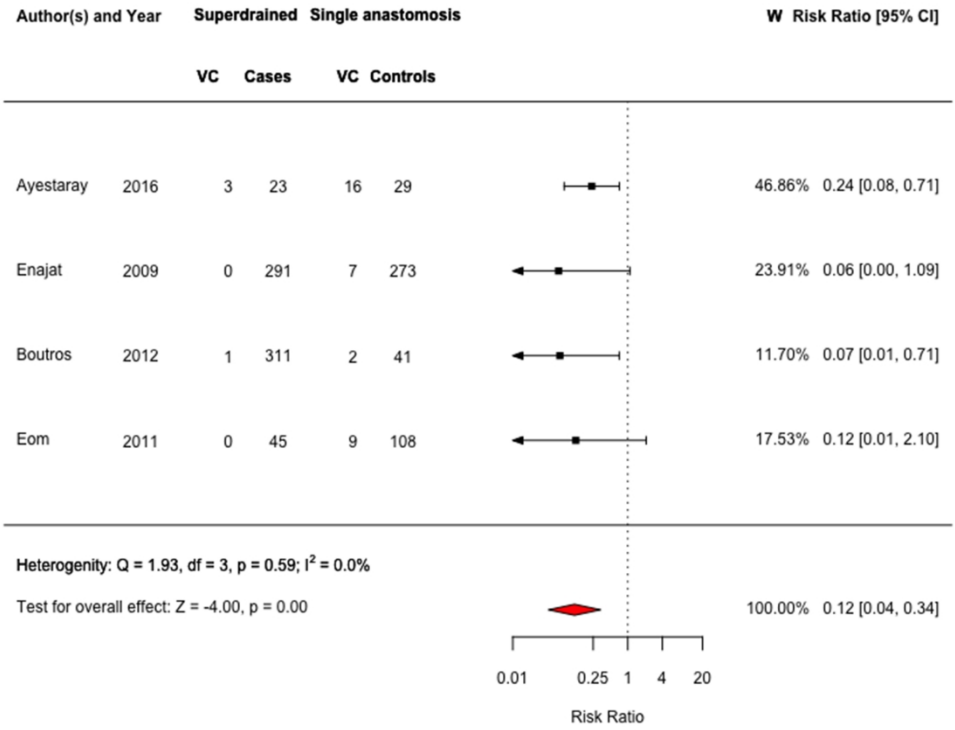


Figure 2

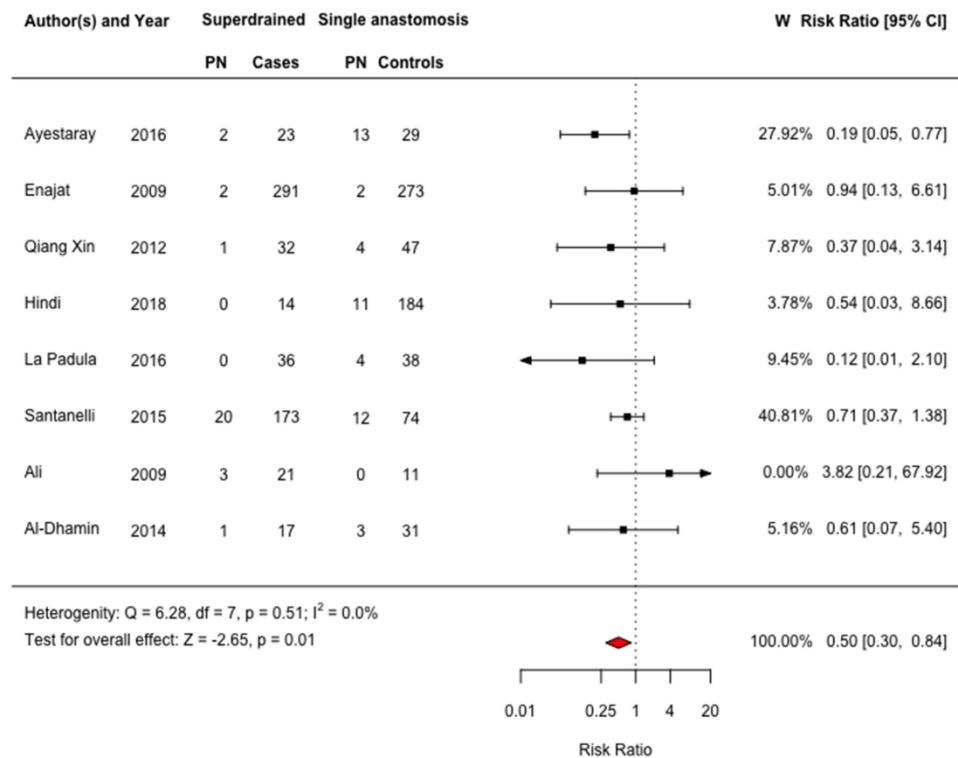


Figure 3

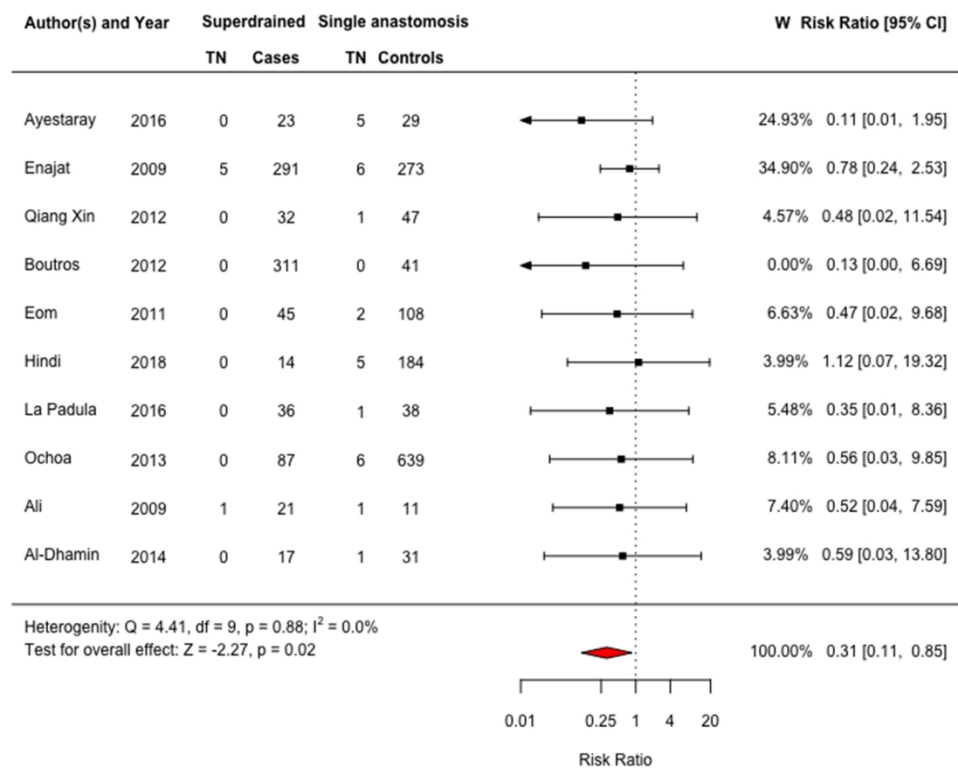


Figure 4

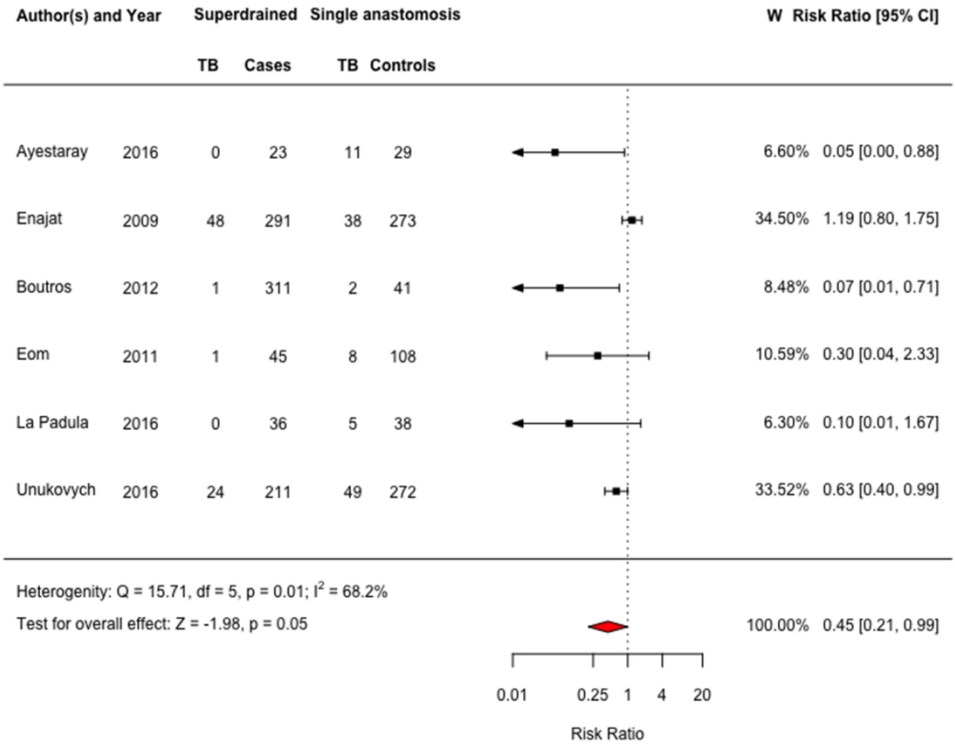


Figure 5

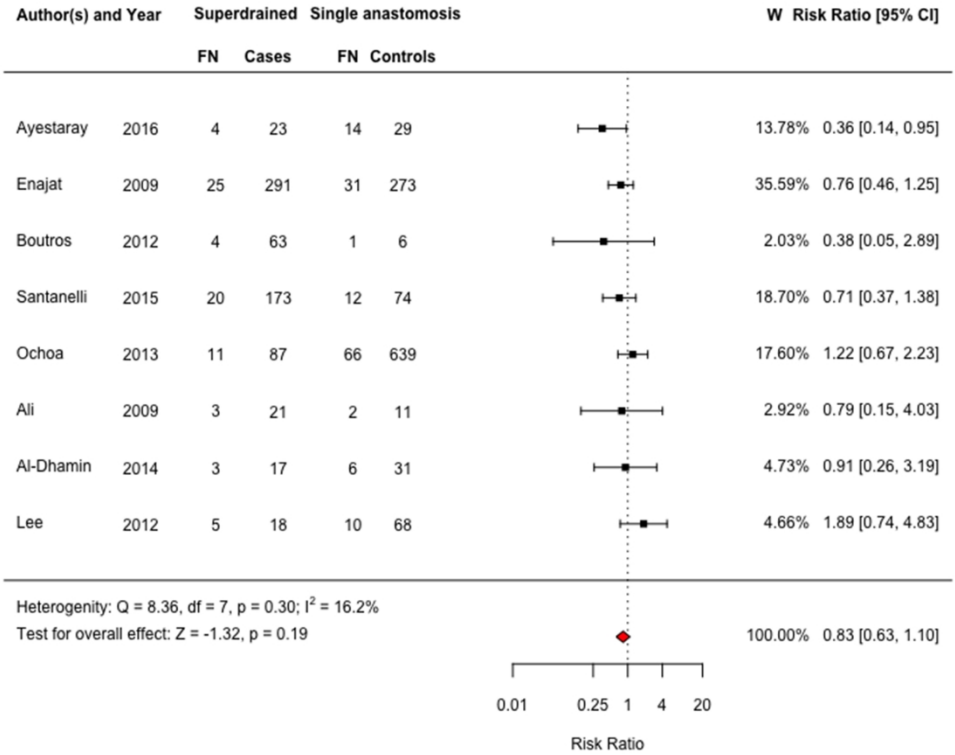


Figure 6

TABLE 1. Studies included in our meta-analysis

Study	Study type	Flap 2° vein	Recipient vein	N. cases	N. controls	Mean OT cases (minutes)	Mean OT controls (minutes)	Difference in OT (minutes)
Ali, 2010	Retrospective cohort	SIEV	IMV	21	11	Na	Na	Na
Enajat, 2010	Retrospective cohort	SIEV	Cephalic vein	291	273	383	385	- 2
Eom, 2011	Retrospective cohort	SIEV	LTV, branch of the TAV, IMVp	45	108	Na	Na	Na
Lee, 2012	Retrospective cohort	SIEV	Na	18	68	Na	Na	Na
Xin, 2012	Retrospective cohort	SIEV	TDV, LTV, ICV, SIEV, DIEV	32	47	396	366	+ 30
Boutros, 2013	Retrospective cohort	SIEV	IMVp, IMV	311	41	Na	Na	Na
Ochoa, 2013	Retrospective cohort	SIEV	IMV	87	639	314	253	+ 61
Al-Dhamin, 2014	Retrospective cohort	SIEV	IMVr	31	17	Na	Na	Na
Santanelli, 2015	Retrospective cohort	SIEV	SCV, TDV, IMV, SUV, LTV	173	74	Na	Na	Na
Ayestaray, 2016	Prospective cohort, randomized	SIEV	TAV	23	29	510	405	+ 105
La Padula, 2016	Retrospective cohort	SIEV or 2 nd DIEV	IMVr	36	38	Na	Na	Na
Unukovych, 2016	Retrospective cohort	SIEV	Na	211	272	Na	Na	Na
Al Hindi, 2019	Retrospective cohort	SIEV	Cephalic vein	14	184	461	356	+ 105

1 OT: operating time, Na: not available, TAV: thoracoacromial vein, TDV: thoracodorsal vein, ICV: intercostal vein, IMV: internal mammary vein, LTV: lateral
2 thoracic vein, IMVr: internal mammary vein retrograde branch, IMVp: internal mammary vein perforator, SCV: scapular circumflex vein, SUV: subscapular
3 vein, AV: axillary vein
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5 “Cases”: flaps with a second venous anastomosis. “Controls”: flaps with a single venous anastomosis
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7 In bold are the 7 studies that were not analyzed in the previous published metanalysis by Lee, 2017
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	Lee and Mun, 2017	Pignatti et al, 2020
<i>Articles included</i>	Enajat, 2010 Lee, 2012 Xin, 2012 Santanelli, 2015 Boutros, 2013 Al-Dhamin, 2014	Ali, 2010 Enajat, 2010 Eom, 2011 Lee, 2012 Xin, 2012 Boutros, 2013 Ochoa, 2013 Al-Dhamin, 2014 Santanelli, 2015 Ayestaray, 2016 La Padula, 2016 Unukovich, 2016 Al Hindi, 2019
<i>Statistical Analysis</i>	Pooled RRs and corresponding 95% confidence intervals (CI) for perfusion-related complications were calculated by the Mantel–Haenszel test in RevMan 5.3. The heterogeneity among studies was evaluated with the I2 test. When I2 ranged from 0 to 30% a fixed- effects model was used. When the value of I2 exceeded 30%, a random effect model was used.	The Mantel-Haenszel test in R was used to calculate Pooled RRs and corresponding 95% confidence intervals (CI) for perfusion-related complications. The heterogeneity among studies was analyzed by using the I2 test. We used a fixed-effects model for I2 ranging from 0 to 30%; in the case of I2> 30%, we applied a random effect model using the DerSimonian & Liard method.
<i>N. of flaps. TOTAL</i> <i>- N. of cases</i> <i>- N. of controls</i>	1376 - 842 - 534	3094 - 1279 - 1815
<i>Total flap loss</i>	RR: 0.97, 95% CI: 0.36-2.57; p = 0.94	RR: 0.31, 95% CI: 0.11-0.85; p = 0.023
<i>Partial flap lost</i>	RR: 0.59, 95% CI: 0.18-1.94; p = 0.39	RR: 0.50, 95% CI: 0.30-0.84; p = 0.008
<i>Fat necrosis</i>	RR: 0.87, 95% CI: 0.58-1.30; p = 0.49	RR: 0.83, 95% CI: 0.63-1.10; p = 0.19
<i>Venous congestion</i>	RR: 0.06, 95% CI: 0.01-0.51; p = 0.01	RR: 0.12, 95% CI: 0.04-0.34; p < 0,001
<i>Conclusion</i>	“The present review demonstrated that superdrainage using SIEV reduces the risk of flap congestion notably, while having little influence on flap survival. With regard to partial flap necrosis including partial flap loss and fat necrosis, general trends toward decreased risks were observed.	“In conclusion, our meta-analysis shows, that venous super-drainage, i.e. performing a second venous anastomosis between the superficial venous system and a recipient vein, provides a statistical advantage in terms of venous congestion and related complications in DIEP flaps

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	However, statistical significance was not achieved and further studies would be needed."	for breast reconstruction."
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TABLE 2. Comparison between our meta-analysis and the one from Lee and Mun

RR: Risk Ratio. Cases: flaps with a second venous anastomosis. Controls: only one venous anastomosis

The 7 studies that were not analyzed in the previous published metanalysis by Lee, 2017 appear in **bold**.

In the fourth row, the results with statistical evidence appear in **bold**.