Anaesthetic implications of free-flap microvascular surgery for head and neck malignancies – A relook

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**Abstract**

Head and neck free-flap microvascular surgery is a type of surgery where multiple anaesthetic factors play a very important role in the outcome of the surgery while the conduct of anaesthesia itself may be quite challenging for the anaesthesiologist. In microvascular reconstruction of head and neck malignancies, flaps are used to reconstruct a primary defect formed by wide local excision. A free flap is raised after removing the neurovascular pedicle from the donor site and transplanting it by microvascular anastomosis to the new location. This gives rise to a secondary defect which is then repaired by direct suture or skin graft. The anaesthesiologist’s role includes optimizing the physiological conditions for the survival of the flap while decreasing morbidity. These patients have a mortality rate of 2.1%, median length of stay 11 days and a flap survival rate of 95%. Failure rate of free flaps is approximately 4% with a rate of re-exploration about 10%. This is an attempt to highlight the numerous causes of free-flap failure along with discussion of the anaesthesia-related issues that are faced during this type of surgery. The various pre-, intra- and postoperative factors affecting flap survival and overall postoperative outcome in the patient are discussed here.

**Keywords:** Anaesthetic factors, flap survival, free-flap, head and neck malignancies, microvascular

**Introduction**

Proper conduct of anaesthesia is required for good surgical outcome in any surgery. Multiple anaesthetic factors play a very important role in the outcome of free-flap microvascular surgery for head and neck malignancies while the conduct of anaesthesia itself may be challenging for the anaesthesiologist due to difficult airway, massive resection and long duration surgery. The anaesthesiologist’s role includes optimizing the physiological conditions for the survival of the flap while decreasing morbidity. These patients have a mortality rate of 2.1%, median length of stay 11 days and a flap survival rate of 95%. Failure rate of free flaps is approximately 4% with a rate of re-exploration about 10%. This is an attempt to highlight the numerous causes of free-flap failure along with discussion of the anaesthesia-related issues that are faced during this type of surgery. Search was carried out using a combination of keywords such as microvascular surgery, free-flap surgery, head and neck reconstruction, anaesthetic factors, hypothermia, steroids, vasopressors, duration of surgery, preanesthetic checkup, free flap thrombosis, free flap necrosis and difficult airway, from various databases such as Pubmed, Cochrane and Google Scholar.

**Surgical Procedure**

A free flap is raised after removing the neurovascular pedicle from the donor site and transplanted by microvascular anastomosis to a new location. Whereas the flaps are used to reconstruct a primary defect formed by wide local excision around the head and neck malignancy, it gives rise to a secondary defect which is then repaired. Various donor sites include radial/ulnar forearm, latissimus dorsi, rectus...
abdominis muscle and groin flap.\textsuperscript{[3]} The preference for donor sites changed significantly over the last decade. For example, the fibula came to be preferred over the iliac crest for mandibular reconstructions as it can be harvested easily under tourniquet and its cutaneous unit used for lining the aerodigestive tract and/or for skin coverage.\textsuperscript{[2]}

### Causes of Free-Flap Failure

Numerous causes of free flap failure that have been cited include arteriovenous thrombosis, vasospasm; mechanical compression due to dressings or positioning; inadequate surgical anastomosis, insufficient venous drainage, flap edema due to excessive use of crystalloids/haemodilution, histamine release; excessive flap manipulation; generalized vasoconstriction due to hypovolemia, hypothermia, pain, respiratory alkalosis; myocardial depressor drugs (anaesthetics, Ca2+ channel blockers). Various anaesthesia-related factors may affect both central haemodynamic stability and regional blood flow.\textsuperscript{[1,3,4]} Poor blood flow to the flap is cited to be the primary cause of flap failure as per evidence.\textsuperscript{[5]} Way back in 1985, Aps et al.\textsuperscript{[6]} reported that local vasospasm could be prevented by the use of direct arteriolar vasodilators (Sodium Nitroprusside). The various factors affecting flap survival and overall post-operative outcome in the patient are discussed below [Table 1].

### Preoperative factors
Age per se has no direct effect on free flap outcome but age over 55 years may increase medical complications following surgery.\textsuperscript{[2]}

Higher American Society of Anaesthesiologists (ASA) class, cigarette smoking and weight loss more than 10% before surgery have been shown to be associated with flap failure.\textsuperscript{[2,7,8]} Preoperative comorbidity did not have any statistically significant effect but it did seem to increase the absolute risk more than four-fold.\textsuperscript{[2]}

Other factors such as sex, chemotherapy, radiotherapy, stage IV cancer, tobacco use, and preoperative use of blood products have been shown not to correlate with postoperative complications.\textsuperscript{[9]}

Even though some studies have shown a negative effect of irradiation due to its adverse effect on blood vessels and delayed healing,\textsuperscript{[2]} others reported no association with either flap outcome or medical complications.\textsuperscript{[10]} Preoperative radiation may affect the airway however.\textsuperscript{[11]}

A study involving 2,846 patients with head and neck cancer found diabetes mellitus, peripheral vascular disease, renal failure, preoperative radiotherapy to be significant predictors.\textsuperscript{[12]}

### Intraoperative factors

#### Type of anaesthesia/anaesthetic drugs
Balanced anaesthesia with benzodiazepines and opioids along with inhalational or intravenous agents are used across centres. Air/oxygen combination may be used with a volatile agent or Total Intravenous Anaesthesia (TIVA).\textsuperscript{[3]} Induction with sevoflurane is preferred in cases where a difficult intubation is anticipated as spontaneous ventilation can be preserved for managing the airway. In 2016, a randomised controlled trial [RCT] comparing balanced anaesthesia with sevoflurane and TIVA (propofol/remifentanil) monitored regional tissue oximetry to conclude that sevoflurane reduces ischemia-reperfusion injury.\textsuperscript{[13]} However, a recent study by Chang et al.\textsuperscript{[14]} comparing TIVA with inhalation anaesthesia (sevoflurane/desflurane) observed that the TIVA group required less perioperative fluids and developed fewer pulmonary complications. Infusion of remifentanil, a short-acting opioid, provides excellent intra-operative analgesia, rapid control of blood pressure, marked vasodilatation and also obviates the need for a muscle relaxant.\textsuperscript{[3]}

| Table 1: Perioperative parameters affecting flap outcome |
|--------------------------------------------------------|
| **Preoperative factors**                               |
| Age                                                    |
| Sex                                                    |
| ASA status                                             |
| Smoking/tobacco abuse                                  |
| Preoperative use of blood products                     |
| Preoperative haemoglobin                               |
| Preoperative haematocrit                               |
| Preoperative comorbidity e.g., Diabetes Mellitus,       |
| peripheral vascular disease, renal failure             |
| Preoperative Chemotherapy                              |
| Preoperative Radiotherapy                              |
| Weight loss >10%                                       |
| **Intraoperative factors**                             |
| Type of Anaesthesia/anaesthetic drugs                  |
| Hypothermia                                            |
| Hypoxia (\textit{SaO}_2 >94\%)                         |
| Haematocrit                                            |
| Hypotension                                            |
| Goal-directed fluid therapy                            |
| Vasoactive drugs                                       |
| Antithrombotic therapy                                 |
| Duration of surgery                                    |
| **Postoperative factors**                              |
| Hypothermia                                            |
| Hypoxia (\textit{SaO}_2 >94\%)                         |
| Haematocrit                                            |
| Hypotension                                            |
| Postoperative bleeding/edema in flap                   |
| Urine output                                           |
| Airway complications                                   |
| Pulmonary complications, e.g., pulmonary edema,         |
| pneumonia,atelectasis                                  |

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Intravenous dexmedetomidine may be continued as an infusion for up to 12 hours in the ICU postoperatively. The reservation against the use of this α2 agonist in free flap surgeries is that it could cause vasoconstriction leading to flap failure. It has however been found that dexmedetomidine maintains postoperative hemodynamics without any increase in flap compromise. It also decreases postoperative agitation.[15,16]

After induction of anaesthesia the trachea has to be intubated with a cuffed endotracheal tube (ETT) followed by careful packing around the tube to prevent aspiration of blood. Controlled ventilation is then maintained while avoiding both hyperoxia and hypocarbia as they can trigger arteriolar vasoconstriction.[6]

Analgesia in free flap harvesting site may be achieved with regional anaesthesia techniques like epidural or local perineural catheter. The evidence regarding the influence of regional anaesthesia on microvascular free flap surgery is inconclusive. Sympathetic blockade with regional anaesthesia has been thought to cause vasodilatation leading to better flap survival but was not helpful in preventing vasoformdue to surgical manipulation whereas sodium nitroprusside infusion was. Also the sympathetically denervated neo-vascularized tissue may be adversely affected by a steal phenomenon.[16,17] Mini-catheters have been used safely to inject local anaesthetic into the fibula[18] and abdominal donor site with resultant decrease in opioid and antiemetic use as well as shorter hospital stay.[19]

Continuous paravertebral block at levels T1 and T2 has been shown to improve tissue perfusion in cases of maxillofacial free flap surgery.[20] Heparin (unfractionated) administered during flap harvesting warrants careful monitoring before and after removal of epidural catheter.[21]

Local anaesthesia with monitored anaesthesia care is used for procedures like flap thinning after free flap reconstruction in head and neck malignancies.

Monitoring
Apart from the routine monitoring (electrocardiography-ECG, Non-invasive Blood pressure-NIBP, Pulse Oximetry-SPO2, End tidal carbon dioxide-ETCO2, temperature and neuromuscular monitoring), other monitors like Central venous pressure (CVP) and Invasive arterial blood pressure (IBP) may be required in these cases where huge amount of blood loss may occur. Clinical monitoring of the patient is important with intraoperative urine output, blood loss, glucose and Arterial Blood Gas (ABG) assessment. Postoperatively too the patient has to be carefully monitored for sedation level and pain apart from vitals monitoring. Various studies where goal directed fluid therapy has been followed used arterial pulse contour device for Cardiac output (CO), Cardiac Index (CI) and Stroke Volume (SV) monitoring.[22,23]

Airway management
Being prepared for difficult airway is of utmost necessity. Many a patient may present for surgery post-radiation which further complicates matters due to involvement of various tissues.[11]

Awake fibroptic intubation of the trachea is usually the preferred method. Fibroptic intubation after induction of anaesthesia is another option when there is no risk of loss of the airway after induction. In extremely difficult cases of massive resection followed by reconstruction, elective tracheostomy is desirable. Different types of cuffed ETT that may used are:

- Polycrystalline (PVC) ETT
- Reinforced ETT
- Ring Adair Elwyn (RAE) tubes
- Tracheostomy tubes

Routes of insertion of ETT may be per oral, nasal, submental or retromolar in some cases.

Tissue edema may compromise airway patency postoperatively and mechanical ventilation for a few hours allows edema to subside before extubation. The difficult airway trolley has to be available for extubation. Prophylactic administration of intravenous steroids (dexamethasone, methylprednisolone) reduces the incidence of laryngeal oedema and reintubation rate after extubation in adults.[24-26] Due to the area of surgery, peritubal leak may not be easy to assess. Use of the airway exchange catheter (AEC) may aid tracheal reintubation in these patients.[27]

Intravenous access and fluid management
Wide bore intravenous access is a must for these patients. The limb from where free flap is to be harvested should be marked preoperatively to avoid securing venous access in this arm.

Central venous catheter (mostly subclavian vein) should be inserted in these patients for guidance in fluid resuscitation. Goal-directed fluid therapy using minimally invasive cardiac output monitoring could improve haemodynamics which in turn would lead to less fluid administration during the perioperative period. Cardiac output measurement, Cardiac Index (CI), Stroke volume index (SVI) may be monitored intraoperatively with the arterial pulse contour device.[22] No significant difference in patient outcome was seen when goal-directed fluid management (GDFM) was followed in another RCT with Stroke Volume (SV) monitoring. GDFM however led to a decrease in the overall
volume of crystalloids infused while increasing the volume of colloids.[23]

Exaggerated tissue edema due to low colloid oncotic pressure after crystalloid use of more than 7 L may worsen edema due to tissue handling.[2,6] Huge blood loss may warrant blood transfusion in many cases with its inherent complications. A postoperative transfusion trigger of haematocrit less than 25 percent has been shown to decrease the rate of blood transfusion while decreasing flap failure rates.[28]

Vasoactive drugs
Vasopressors used during anaesthesia have been traditionally thought to be one of the causes of poor blood flow to the flap.[29] Other studies however have shown that the type (dopamine, noradrenaline and metaraminol) or method of usage of vasoconstrictors could not be associated to flap failure directly and in fact vasoconstrictors such as noradrenaline have been shown to be beneficial by maintaining blood flow to the flap.[5,30,31] Sodium nitroprusside improves blood flow in the free flap on direct administration onto the vessels being anastomosed.[6] It has the advantage over alpha blocker (labetolol), of fast and profound vasodilation in the sympathetically denervated free flap without undue effect on the sympathetic nervous system.[32]

Antithrombotic therapy
Some agents that are used with varied protocols to decrease platelet functioning, improve blood flow or decrease blood viscosity are Aspirin, Dextran 40 and Unfractionated Heparin. Statins have vasoprotective and anti-inflammatory actions but their role in preventing anastomotic thrombosis is not known.[33]

Dextran infusion has traditionally been used to improve perfusion in the free flap vessels.[33,34] However, the routine use of dextran has been questioned due to adverse effects such as anaphylactoid reactions, adult respiratory distress syndrome, cardiac overload, haemorrhage, and renal damage. The patency rates of free flap reconstructions and thrombotic complications were not statistically different from patients not receiving dextran infusion.[35,36]

Once a thrombus is formed, options that are available are mechanical re-exploration, thrombolysis with agents like tPA.[37]

Temperature control
Long duration of surgery under GA, multiple exposure sites to cool OR temperatures and cool intravenous fluids often lead to hypothermia. Hypothermia has been implicated in causing various postoperative complications like partial or complete flap loss, delayed wound healing, local vasoconstriction, increased viscosity/haematocrit, coagulopathy leading to haematoma formation and infectious complications. Postoperative shivering may lead to increased oxygen consumption, hypoxia, arrhythmias and myocardial events.[1,9,38,40]

Surgical factors
Operative time has been shown to be positively associated with postoperative complications, morbidity, and prolonged length of stay in various type of surgeries.[41,42] A study on 2,008 patients[7] observed a correlation between operative time and the incidence of early flap failure and postoperative complications following microvascular tissue transfer surgery using the American College of Surgeons National Surgical Quality Improvement Program (ACS-NSQIP) database. Other studies have however observed that duration of surgery did not per se influence the medical or surgical outcomes.[2]

The number of surgeons may affect the outcome which has been shown to improve when preoperative selection and postoperative management is handled by one person.[2] Another study has however reported same outcome even with multiple surgeons.[43]

Postoperative factors
Postoperative care should include maintaining normothermia, normal blood pressure, haematocrit around 30%, urine output >1 ml kg⁻¹ h⁻¹, SaO₂ >94% (oxygen for the first 24 h), regular inspection of the flap and continuous monitoring of blood flow in the flap by temperature and laser Doppler.[3] Edema and bleeding in the flap site may lead to airway obstruction. Common modalities of postoperative pain management are intravenous non-steroidal anti-inflammatory drug (NSAID), paracetamol, and opioids. Mini-catheters have been used safely to inject local anaesthetic into the fibular donor site after flap harvesting for reconstruction of the head-and-neck area with successful outcome.[18] Pulmonary complications such as pulmonary edema, pneumonia or atelectasis may occur in these patients.[14]

Grafts should be monitored for 24 hours for arterial spasm, graft oedema and venous occlusion. Re-exploration of the graft may be required if signs of ischaemia occur. Graft oedema can be reduced by elevation of the recipient site, IV drugs like single dose of dexamethasone (40 mg) and mannitol 10% (0.5 g/kg).[6] Routine use of IV heparin and dextran infusion are not followed by many centres due to concern over rebleed.[6,36]

Conclusion
Microvascular flap surgery for head and neck malignancies is a challenge to the anaesthesiologist and optimization of the physiologic conditions for flap survival while keeping
morbidity at check is the goal. Knowledge of the various aspects of this surgery along with close communication with the surgeon and regulation of the numerous factors that may affect flap survival are imperative to ensure a favourable outcome.

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**Conflicts of interest**
There are no conflicts of interest.

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