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# Quarterly Report on Research Activities in the Adult Neurosurgery Program

January 1st – March 31st 2024

Prepared by

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March 31st, 2024

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# 1. INTRODUCTION

This report contains up to date information on the ongoing research projects that are supported by the Clinical Research Coordinators (CRC) of the University of British Columbia's (UBC's) Division of Neurosurgery at the Clinical Academic Campus of Vancouver General Hospital (VGH) for the period of October 1<sup>st</sup> to December 31<sup>st</sup>, 2023. The main objective of the report is to familiarize the staff of the Division of Neurosurgery of UBC with the current research activities that are being supported by their CRCs. The studies that are supported by the CRC in this report are divided into two categories of ongoing studies: prospective studies, and retrospective studies. The number of studies per category is presented in the table below.

Number of Ongoing Studies								
Prospective Retrospective Inactive or Complete Total Studies								
12 17 4 33								

Detailed description of the purpose, objective, budget and sample size of each study supported by the CRC is presented in the next four sections of this report.

\*\*This report does not encompass research projects in the Division's pediatric neurosurgery, functional neurosurgery and spine neurosurgery programs.

# 2. ONGOING PROSPECTIVE STUDIES

#### 1. <u>Quality of Life in Patients Diagnosed with Moyamova Disease: Cross-Sectional</u> <u>Study:</u>

Funding	Source	Study period	Anticipate d Enrolment	# of subjects enrolled	Approvals	Status	Abstract/P aper/Manu script
Yes	Rare Diseases Foundatio n	May 2017 - Oct 2025	100	71	Yes	Approved	N/A

PI: Dr. Gooderham; Co-PI: Drs. Dandurand, Yip

#### Purpose

Moyamoya disease is a rare and chronic disease characterized by the progressive occlusion of intracranial vessels. The supraclinoid carotid arteries are the first arteries affected. It rarely affects the posterior circulation. Small collateral vessels begin to form at the base of the brain as the larger vessels become occluded giving it the characteristic appearance of a «puff of smoke» on angiographic radiological studies. This disease can present with ischemic stroke or intracranial hemorrhage depending on the age of the patient.

#### Objectives

The main goal of the present study is to identify how does the diagnosis of Moyamoya disease, its different clinical presentations and its subsequent treatment impact quality of life as measured by SF-36, EQ5D and 49-item Stroke-Specific Quality of Life Scale (SSQOL) in patients. We aim to verify if the choice of technique (direct or indirect bypass) has an impact on quality of life. We will explore other variables such as clinical presentation (incidental, ischemic or hemorrhagic), radiological features (cerebrovascular reserve and evidence of ischemic stroke), gender, medical comorbidities, postoperative complications and length of time after diagnosis and treatment.

There are 71 participants currently enrolled in the study. 5 were enrolled in the last quartile

Funding	Source	Study period	Anticipated Enrolment	# of subjects enrolled	Approvals	Status	Abstract/P aper/Manu script
N/A	N/A	July 2022 -Jun 2025	50	12	Yes	Active	N/A

#### 2. <u>Timing of Mobilization After Burr Hole Drainage of cSDH: a randomized study:</u> PI: Dr. Rvoio Akagami: Co-I: Drs. Chang, Craig, Rizzuto, Rebchuk, Houniet, Wang

### Purpose

This is a two treatment arm, randomized, prospective study to minimize re-do burr-hole drainage procedures and any other associated complications in patients with chronic subdural

haematomas.

Primary Outcomes:

- Recurrence requiring re-do drainage within the 1st month post-operatively
- Recurrence requiring re-do drainage between the 1st and 3rd months post-operatively

The timing of when to mobilize patients after burr-hole drainage of chronic subdural haematomas remains controversial. Traditionally, patients have been subjected to delayed mobilization in order to allow for the theoretical re-expansion of the brain and to decrease recurrence. Timing of bed rest is not consistent among centres and varies from immediately after to 7 days after surgery.

# Objective

The objective is to determine optimal timing of mobilization in CSDH patients following a burrhole drainage.

There are 12 participants currently enrolled in the study, 0 participants were enrolled during the last quarter.

#### 3. <u>Adult Hydrocephalus Clinical Research Network (AHCRN):</u> PI: Dr. Thomas Zwimpfer

Funding	Source	Study period	Anticipated Enrolment	# of subjects enrolled	Approvals	Status	Abstract/P aper/Manu script
N/A	N/A	Nov 2014 -	Perpetual	651	Yes	Active	N/A

A multi-centre and multinational registry that collects data on adult hydrocephalus patients to characterize the etiology, understand variability, progression, and current treatment practices for hydrocephalus patients.

The overall purpose of the Registry is to establish and maintain a hydrocephalus patient event database for the Clinical Centres of the AHCRN, a research network newly established to investigate clinical management of adult hydrocephalus.

# Objectives

- To describe the natural history and treatment response for adults with previously untreated congenital hydrocephalus
- To describe the assessment and treatment of patients with Normal Pressure Hydrocephalus (NPH)
- To describe the complications associated with shunt surgery
- To determine the role for treatment with Endoscopic Third Ventriculostomy (ETV)

The Registry will provide previously unavailable epidemiological information about hydrocephalus patients seen throughout the participating Clinical Centers. This information will provide the basis for multi-institutional studies to be carried out by the AHCRN that may ultimately improve the clinical care for adults with hydrocephalus throughout the world. The continuing collection of such information serves to provide data necessary for hypothesis generation and study design. Examples of preliminary study designs include, but are not limited to, the following: preliminary power analysis, sample size determination, and recruitment projections. Radiologic imaging data will provide a unique opportunity to assess aspects of adult hydrocephalus diagnosis, management, and outcomes.

# There are 2,244 participants enrolled in the AHCRN Registry at all participating sites. Of those, 651 are from the VGH site with 19 patients enrolled in the last quarter.

# 4. <u>Neuroscience of the Human Brain in Health and Disorder:</u>

**PI:** Dr. Mark Cembrowski (UBC Dept. Cellular and Physiological Sciences); **Co-I's**: Dr. Redekop, Fatehi, Hirsch-Reinshagen

Funding	Source	Study period	Anticipated Enrolment	# of subjects enrolled	Approvals	Status	Abstract/P aper/Manu script
Yes	Grant - Internal Funds	Nov 2020 -	120	31	Yes	Active	N/A

(1) Understand organization and function of the healthy (i.e., pathologically unremarkable) human brain at molecular, cellular, and circuit levels; (2) use this understanding of the healthy brain to interpret dysregulation during epilepsy.

# Objectives

(1) Characterize the molecular, cellular, and circuit properties of surgically resected discard human brain tissue in healthy cortex, (2) Compare this to analogous characterizations in brain tissue from epileptic foci to identify molecular, cellular, and circuit dysregulation in epilepsy.

### **Research design**

Non-diagnostic discard brain tissue from Dr. Redekop and his staff, obtained via standard surgical resections from informed consenting participants, will be de-identified by Dr. Redekop or a member of VGH research staff, and via Dr. Hirsch- Reinshagen and VGH pathology, received by Dr. Cembrowski to be used in neuroscientific research examining molecular, cellular, and circuit properties. Acute experiments will involve:

- transcriptomics and epigenomics (tissue from n=12 participants needed for each of epileptic and non-epileptic datasets, in order to acquire sufficient statistical power based upon previously published results)
- whole-cell patch-clamp electrophysiology and morphological reconstructions (n=50 neurons needed for each of epileptic and non-epileptic datasets, likely requiring tissue from ~10 participants, in order to acquire sufficient statistical power based upon previously published results)

There are 31 participants currently enrolled in this study. 4 were enrolled this quarter.

# 5. <u>A Placebo-Controlled Efficacy in iNPH Shunting (PENS) Trial:</u>

PI: Dr. Thomas Zwimpfer

Funding	Source	Study period	Anticipated Enrolment at VGH Site	# of subjects enrolled	Approval	Status	Abstract/Paper/ Manuscript
Yes	National Institutes of Health	Nov 2022 - Jan 2028	10	4	Yes	Active	N/A

Although idiopathic normal pressure hydrocephalus (INPH) has been recognized for five decades, barriers still exist in recognition, referral and accurate diagnosis. Hesitance in referring elderly patients for surgical treatment of INPH results from an incomplete understanding of its pathophysiology, controversy over the appropriate diagnostic work up, and a significant concern about the effectiveness and complications of surgical treatment. The lack of consensus regarding tests predicting outcome of surgery in INPH, and the skepticism of INPH in the neurology and neurosurgery communities reflect the limitations of INPH clinical research to support current INPH practices. Convincing proof of shunting effectiveness is likely to increase the number of INPH patients getting adequate treatment. The Placebo-Controlled Effectiveness in INPH Shunting (PENS) trial is a multi- center blinded, randomized, placebo-controlled design investigation of cerebrospinal fluid (CSF) shunt surgery.

#### Objective

### Primary Objective:

The primary objective is the evaluation of CSF shunting in iNPH participants through a randomized comparison of improvement from baseline at three months between active (open shunt) and placebo-controlled (closed shunt) groups, using the primary endpoint of gait velocity to test the primary hypothesis: the treatment of idiopathic normal pressure hydrocephalus (iNPH) with an open shunt results in improved gait velocity.

Secondary Objectives:

- 1. Evaluate the effect of shunting on improving gait and balance between active and placebo-controlled groups at three months using the Tinetti assessment
- 2. Evaluate the effect of shunting on improving global cognition between active and placebo-controlled groups at three months using the total Montreal Cognitive Assessment (MoCA) score
- 3. Evaluate the effect of shunting on bladder control between active and placebo-controlled groups at three months using the OAB-q sf.

#### Justification

There are 5 major manufacturers of shunts. Until now the valves were either differential pressure valves or flow-regulated valves and none of them had the ability to be turned off even if clinically indicated. Thus, doing a placebo study of shunts often involved tying a ligature in the shunt catheter with variable results and adding complexity and additional intervention to untie the ligature. With the release of the new Codman Certas Plus 2.0, a virtual off setting is now available to stop flow of CSF through the shunt system unless intracranial pressures exceed 400 mm which has not been documented in participants with iNPH. This shunt would also rapidly enable lowering settings if indicated in the judgement of the treating physician in the placebo

arm without necessitating invasive intervention. No other commercially available shunt appropriate for treating NPH offers these features.

There are 100 participants anticipated to be enrolled across 20 sites. 10 participants will be enrolled at the Vancouver site. 0 participants were enrolled in the last quarter.

# 6. <u>Aneurysmal Subarachnoid Hemorrhage - Red Blood Cell Transfusion and Outcome</u> (SAHaRA): A Randomized Controlled Trial:

Funding	Source	Study period	# of subject enrolled	Approvals	Status
Grant	CIHR	2017-2022	29	Yes	Recruitment ongoing

#### Purpose

We propose a multicenter pragmatic randomized trial in patients with aSAH that will compare the effect of a liberal to a restrictive RBC transfusion strategy on the combined rate of death and severe disability at 12 months.

#### Hypothesis

We hypothesize that in adult patients suffering from aSAH and anemia, a liberal RBC transfusion strategy as compared to a restrictive RBC transfusion strategy decreases the combined rate of death and severe disability at 12 months (using the modified Rankin Scale)

#### Justification

Aneurysmal subarachnoid hemorrhage (aSAH) is a devastating illness caused by the spontaneous rupture of an enlarged, weakened artery in the brain. It affects a young population and is a significant cause of premature death and loss of potential life years, similar in magnitude to ischemic stroke. It is a common neurologic reason for intensive care unit (ICU) admission and is associated with a mortality rate of 35% in North America. Less than one third of afflicted patients make a full recovery and 20% of survivors experience significant morbidity and impacts on daily living.

# *There are 478 participants enrolled across 20 sites. 29 participants have been enrolled in Vancouver*

#### 7. Vancouver Ruptured Aneurysm Database (VRAD)

PI: Dr. Peter Gooderham; Co-I's: Drs. Haw, Rebchuk, Chang, Redekop, Rizzuto

Funding	Source	Study period	# of subject enrolled	Approvals	Status
None	N/A	2022-2032	147	Approved	Recruitment and Data Collection ongoing

#### Purpose

The overall purpose of the registry will be to establish and maintain a longitudinal aneurysmal subarachnoid hemorrhage patient event database to investigate clinical management of subarachnoid hemorrhage. Through this data, long term functional outcomes will be characterized.

#### Objectives

- To describe the demographic and natural history of patients with subarachnoid hemorrhage
- To characterize the presentation, assessment, and treatment of patients with subarachnoid hemorrhage
- To characterize both open surgical and endovascular management for subarachnoid hemorrhage
- To describe hospital-related complications and long-term clinical outcomes for subarachnoid hemorrhage patients

#### **Study Design**

This is an observational prospective cohort study. Participants will be recruited by study investigators. They will be flagged upon presentation to hospital and the neurosurgery service. The neurosurgery service is consulted on all cases of suspected aSAH and intracranial aneurysms at Vancouver General Hospital. Data will be extracted from hospital charts and electronic medical records. We will collect basic demographics, past medical history, clinical presentation, treatment, complications, clinical and functional outcomes. Hospital and clinical data will be collected and stored in an online web-based registry constructed using UBC ARC REDCap. *There are 147 participants enrolled from Vancouver General Hospital. 32 participants have been enrolled in the last quarter.* 

# 8. <u>The SmartForceps System: An Intelligent Device for Real-time Measurement of</u> <u>Forces of Tool-tissue Interaction during Surgery, towards Assessment of Surgical</u> <u>Skill and Performance:</u>

PI: Dr. Serge Makarenko Co-I: Dr. Peter Gooderham

Funding	Source	Study period	# of subject enrolled	Approvals	Status
Grant	CIHR	2022	n/a	REB and VCHRI approved	Data Collection

### Purpose

This study aims to trial SmartForceps, a new tool that is the first of its kind, integrated with force sensors for real-time sensing and monitoring of coagulation, dissection and tool-interaction forces during surgery. This includes a force monitoring system, allowing the ability to monitor and record surgical forces during procedure, providing helpful feedback to surgeons.

### Hypothesis

1. The SmartForceps System will differentiate surgeons by their skill level

2. Surgical task specific force profile will be automatically and reliably recognized and created by the machine-learning module.

3. Shortened operating room time in comparison to historical controls.

# Justification

It has been shown using virtual reality simulators that ~50% of surgical errors are related to the use of inappropriate/excessive force. Currently, the ability to apply optimal force during tooltissue interaction in surgery is only mastered through years of hands-on surgical training. This knowledge remains largely qualitative, as none of the commercially available surgical instruments provide real-time measurement of intra-operative surgical forces. When a surgeon is operating or supervising a trainee, there is no objective means of evaluating the amount of force applied, and trauma to delicate tissues such as brain or nerve tissues often cannot be detected until obvious injuries have occurred.

The SmartForceps is the first of its kind, integrated with force sensors for real-time sensing and monitoring of coagulation, dissection and tool-interaction forces during surgery. The prototype included a calibration station, signal conditioning and filtering module, together with a force monitoring system, allowing the ability to monitor and record surgical forces during procedure.

#### 9. <u>Subarachnoid Hemorrhage - Vasospasm Neuromonitoring</u>

PI: Dr. Peter Gooderham; Co-I's: Drs. Griesdale, Rebchuk, Sekhon

Funding	Source	Study period	# of subject enrolled	Approvals	Status
None	N/A	2022	1	Approved	Recruitment and Data Collection

#### Purpose

Currently, detection of delayed cerebral ischemia in patients presenting with subarachnoid hemorrhage is undertaken by scans or clinical exams, which have limitations. This study will assess whether specialized micro-catheter monitors that are inserted into the brain following aneurysmal subarachnoid hemorrhage can allow for early detection of delayed cerebral ischemia.

#### Objectives

- To examine differences in PbtO2 in aneurysmal subarachnoid hemorrhage (aSAH) patients with delayed cerebral ischemia (DCI) vs those without
- To examine differences in cerebral blood flow in aSAH patients with DCI vs those without
- To examine differences in lactate/pyruvate ratio in aSAH patients with DCI vs those without
- To examine differences in SjvO2 in aSAH patients with DCI vs those without
- To correlate PbtO2, cerebral blood flow, lactate/pyruvate ratios during DCI to functional and quality of life outcomes at discharge, 6 months, 12 months, and 2 years post-ictus

#### **Study Design**

This is a prospective interventional study. Participants will be screened and recruited by study investigators and research assistants. They will be flagged upon presentation to hospital and the neurosurgery service. An intraparenchymal catheter will be placed within the sub-cortical white matter of participants to real-time monitoring and assessment of key information such as brain oxygenation, cerebral blood flow, cerebral metabolism, and exchange of oxygen in the cerebral vascular bed. This will allow us to directly explore how the cerebral microvascular changes in response to vasospasm and in patients that develop delayed cerebral ischemia.

#### The first participant was enrolled in the last quarter.

# 10. <u>Prospective registry for adults with Moyamoya disease and Moyamoya syndrome at the Vancouver Stroke Program</u>

Funding	Source	Study period	# of subjects enrolled	Approvals	Status	Abstract/P aper/Manu script
No	n/a	2023-2027	18	Yes	Approved	N/A

PI: Dr. Yip; Co-PI: Drs. Gooderham, Sheldon, Teal

#### Background

Moyamoya disease is a rare and chronic disease characterized by the progressive occlusion of intracranial vessels. The supraclinoid carotid arteries are the first arteries affected. It rarely affects the posterior circulation. Small collateral vessels begin to form at the base of the brain as the larger vessels become occluded giving it the characteristic appearance of a «puff of smoke» on angiographic radiological studies. This disease can present with ischemic stroke or intracranial hemorrhage depending on the age of the patient.

#### **Objectives**

The purpose of this registry is to improve our understanding of the natural history and outcomes of patients living with Moyamoya disease and Moyamoya vasculopathy. There is currently no curative treatment for Moyamoya, however some interventions exist to reduce the risk of ischemic and hemorrhagic complications as well as the onset of other symptoms. Although these interventions are available, there is a lack of consistent information regarding the demographics and natural history of this patient population, and importantly, their long-term outcomes. This registry will allow us to prospectively assess this patient population overtime which is essential for improved understanding, and treatment, of this rare disease.

There are 18 participants enrolled from Vancouver General Hospital. 5 participants have been enrolled in the last quarter.

#### 11. <u>Moyamoya disease in Indigenous Populations in British Columbia: Presentation,</u> <u>treatment, outcomes, and specific challenges they face</u>

Funding	Source	Study period	# of subjects enrolled	Approvals	Status	Abstract/P aper/Manu script
No	n/a	2023-2024	n/a	n/a	Pending CREB Approval	N/A

PI: Dr. Gooderham; Co-PI: Drs. Yip, Teal, Hounjet

#### Background

Moyamoya disease is a rare and chronic disease characterized by the progressive occlusion of intracranial vessels. The supraclinoid carotid arteries are the first arteries affected. It rarely affects the posterior circulation. Small collateral vessels begin to form at the base of the brain as the larger vessels become occluded giving it the characteristic appearance of a «puff of smoke» on angiographic radiological studies. This disease can present with ischemic stroke or intracranial hemorrhage depending on the age of the patient. This disease has been well-characterized in the literature among Korean and Japanese populations. Clinically, there appears to be a disproportionate representation of Indigenous peoples affected by this disease in British Columbia, but this has not been documented in literature.

#### **Objectives**

This study aims to provide a clinical and radiographic description of Moyamoya disease among Indigenous people in British Columbia, while also assessing data on barriers to care and recovery. With an eye to future research, patients will be asked to identify, as they are comfortable and able, potential avenues to future contact with their community for collaborative research regarding traditional knowledge and stories that may pertain to stroke-like presentations in young people.

# 12. <u>Simulating Brain Tumour Resection to Optimize Onco-Functional Balance in Low-Grade Glioma</u>

Funding	Source	Study period	Anticipat ed Enrolme nt	# of subjects enrolled	Approvals	Status	Abstract/Paper/ Manuscript
Yes	Brain Tumour Foundation of Canada	2024-2025	20	0	REB and VCHRI Approved	Patient Recruitment and MRI setup	N/A

PI: Dr. Stefan Lang, Co-PI: Dr. Mostafa Fatehi, Dr. Serge Makarenko

#### Purpose

Cognition is critical for quality of life in low-grade glioma patients. It is known that brain networks are involved in cognition, and surgery can affect these networks. However, we cannot predict if the surgical plan will result in damage to these networks and a decline in cognition. To overcome this, virtual brain models could be used to simulate the effects of a proposed surgery on brain networks before the real surgery occurs. These models can be personalized using patient MRI data. In this project, we aim to assess the use of these virtual models in predicting cognitive outcomes in low-grade glioma patients.

#### Objectives

The objective of the proposed research study is to assess the utility of personalized whole-brain models in the prediction of cognitive outcomes following surgery in LGG patients.

This study received REB and VCHRI approval in the last quarter.

# **3. ONGOING RETROSPECTIVE STUDIES**

1. Wounded Glioma Syndrome: neurologic worsening in patients with Subtotal Resection in High-Grade Gliomas

PI: Dr. Serge Makarenko Co-I: Dr. Michael Rizzuto, Crystal Ma (Year 2 Medical Student)

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Jan 2000 - Dec 2021	Approved	200	Data Analysis	N/A	N/A

#### Purpose

First line treatment for high-grade gliomas is surgery with goals of maximally safe resection, and subsequent radiation and chemotherapy. Surgery serves other purposes aside from removing the malignancy, it alleviates the raised intracranial pressure through reduction of mass effect, and additional cytoreduction. A greater extent of resection generally results in better survival for patients with high-grade gliomas. Although the ideal treatment for high-grade gliomas is gross total resection, this approach is not always feasible owing to the infiltrative quality of these tumors. Moreover, neurosurgeons may choose to pursue a relatively more conservative approach such as subtotal resection rather than gross-total resection to minimize injury to eloquent areas that are important for overall quality of life and survival. A delicate balance is presented to the neurosurgeon, achieving maximal resection in order to improve and prolong survival, while balancing it against morbidity and preserving quality of life. One of the challenges with subtotal resection in the early post operative period, is that the residual tumour tissue can lead to associated morbidity in the form of post-operative edema and swelling, and worsening neurologic function. Our aim is to investigate if tumor remaining at the surgical site after subtotal resection entails post-operative neurological injury and deficit, whether this may be due to inflammation or another physiological process.

#### Hypothesis

We hypothesize that patients with high-grade gliomas who have undergone subtotal resections will experience an increased risk of post-operative neurological worsening within the first 7-10 days post-operatively.

#### Justification

High-grade gliomas are common and devastating brain malignancies for which surgery is the first-line treatment. Subtotal resections result in residual tumor that may be associated with neurological worsening and other post-operative complications. Our aim is to whether subtotal resections result in immediate post-operative neurological deficits, specifically 7-10 days after surgery, identify associated factors, characterize other complications, and report an ideal threshold for resection that minimizes neurological deficits. We anticipate these findings will further our understanding of improving surgical outcomes to optimize patient quality of life. Furthermore, by helping delineate the safest extent of resection, we hope to provide additional guidance for surgeons in decision-making around resection of these tumours.

# 2. Recurrence in WHO Grade 1 meningiomas – is there a pattern of clustering among recurrent Grade 1 meningiomas that could help target radiation therapy?

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Jan 2000 - Dec 2021	Approved	1078	Data Collection	N/A	N/A

PI: Dr. Serge Makarenko Co-I: Dr. Michael Rizzuto

#### Purpose

We aim to elucidate the pattern and location of tumor recurrence to target radiation therapies for recurrent Grade 1 meningiomas

# Hypothesis

We hypothesize that there will be a pattern of geographic and temporal clustering of recurrence in WHO Grade 1 meningiomas that can allow for targeted adjuvant radiotherapy delivery.

#### Justification

Meningiomas are common neoplasms in the population that may lead to morbidity and potential mortality. While the current literature suggests that certain patient and tumor specific factors (including patient sex, type and grade of meningioma, and molecular signature) can predict recurrence, the efficacy of adjuvant RT in preventing and managing recurrent disease is not well characterized. Additionally, determining geographic patterns of tumor recurrence is important in aiding the delivery of RT in patients with recurrent disease. Understanding when, where and how

RT should be delivered in patients diagnosed with WHO Grade 1 meningiomas will allow for a tailored approach to therapy based on geographic location of the initial tumor and recurrence.

#### 3. A Retrospective Review of Treatment Outcomes for Glioblastoma Patients across British Columbia

PI: Dr. Mostafa Fatehi Co-I: Dr. Serge Makarenko, Dr. Michael Rizzuto

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Mar 2023 - Aug 2024	UBC CREB Approved Awaiting VCHRI Approval	300	Data Collection	N/A	Grant

#### Purpose

To establish overall and median survival of a current cohort of patients treated for glioblastoma in BC undergoing standard treatment algorithms

#### Hypothesis

This study will validate the predicted improvement in glioblastoma outcome against actual practice results for the time period of 2016-2022

#### Justification

This study will serve to document whether patients in BC are experiencing improvements in survival predicted by recent literature reports. Deviance or improvements on these predictions will be surveyed to qualitatively ascertain causation. The actual performance of local treatment strategies must be audited and insurances made that real world results meet expectations.

# 4. Utility of Cerebral Angiography in Identifying Cerebrovascular Pathology in CTA-Negative Subarachnoid Hemorrhage

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Mar 2023 - Aug 2024	Approved	200	Data collection	N/A	N/A

PI: Dr. Mostafa Fatehi Co-I: Dr. Michael Rizzuto

#### Purpose

To document how often a DSA study reveals actional pathology in patients with a CTA-negative subarachnoid hemorrhage, and how often this information results in change of management at Vancouver General Hospital.

#### Hypothesis

DSA will reveal actionable cerebrovascular pathology to a greater extent than repeat or delayed CTA in patients presenting initially with a CTA-negative subarachnoid hemorrhage, and no cerebrovascular pathology on index CTA.

#### Justification

There would be clinical utility in understanding how often a DSA identifies a vascular pathology and thus alters the management of a patient. It would help better understand what tests and interventions may be applicable for patients with CTA negative SAH and could be used to better guide the diagnostic management of these patients. Ultimately, this may potentially decrease the risk to patients through unnecessary diagnostic testing, improving their management and workup.

# 5. Patient outcomes of a subtemporal preauricular infratemporal approach with condylar fossa osteotomy for skull base chondrosarcoma resection:

Study period	Anticipated Number of Patients Reviewed	Approval	Status	Abstract/Paper/ Manuscript	Funding
Aug 2022 - Jun 2024	50	Yes	Active	N/A	N/A

PI: Dr. Ryojo Akagami; Co-I's: Seika Taniguchi

# Purpose

To the knowledge of the investigators, there is an absence of reviews that synthesize and analyze the health-related quality of life (HRQoL) outcomes of skull base chondrosarcoma patients who have undergone surgical management using this surgical approach. This study aims to fill this gap in the literature and provide greater insight into the HRQoL of chondrosarcoma patients. Findings from this study may guide clinicians in the management and integration of a biopsychosocial approach to patient care.

# **Objectives**

This retrospective chart review aims to:

- 1. Evaluate the surgical and clinical outcomes of skull base chondrosarcoma resection using a subtemporal preauricular infratemporal approach with condylar fossa osteotomy.
- 2. Analyze the impacts of skull base chondrosarcoma on overall patient health related quality of life. This would enrich the understanding of the nuanced, biopsychosocial impact of this condition on patient quality of life and potentially guide future long-term patient management.

# **Research Design**

A retrospective chart review of all patients with skull base chondrosarcoma will be first conducted using existing patient data from charts available on Plexia EMR and VGH hospital records/PCIS. Inclusion and exclusion criteria outlined above will be applied to select for retrospective chart analysis. Relevant surgical and clinical parameters that will be collected and reported on will correspond to pre-, peri- and postoperative records (see attached data collection sheet). The SF-36 questionnaire is part of existing postoperative follow up protocol. Answers from these questionnaires will be tabulated and added to the data collected from the retrospective chart review.

# 6. Does temporomandibular joint disruption in performing a condylar osteotomy affect postoperative patient quality of life in skull base tumours:

Study period	Anticipated Number of Patients Reviewed	Approval	Status	Abstract/Paper/ Manuscript	Funding
Oct 2022 - Jun 2024	50	Yes	Active	N/A	N/A

PI: Dr. Ryojo Akagami; Co-I's: Seika Taniguchi

### Purpose

To date, there remains an absence of reviews that evaluates and synthesizes the oral healthspecific quality of life (OHRQoL) following TMJ disruption in the SPI approach to managing chondrosarcomas. Temporomandibular disorders (TMD) are significant public health problems and are the second most common musculoskeletal condition. second to chronic low back pain. The study aims to provide greater insight into the (OHRQoL) of patients with skull-based tumours, namely chondrosarcomas and trigeminal schwannomas.

# Objectives

- 1. Compare the temporomandibular joint specific oral health related quality of life (OHRQoL) between patients who underwent skull base tumor resection (chondrosarcoma or trigeminal schwannoma) with either temporomandibular joint preserving approach or temporomandibular joint disrupting approach as a result of a condylar osteotomy.
- 2. Evaluate the subtemporal preauricular infratemporal approach for chondrosarcomas in relation to post surgical temporomandibular joint complications and overall OHRQoL.

# **Research Design**

The skull base tumors used for comparison will be chondrosarcomas and trigeminal schwannomas; chondrosarcoma resection typically involves a condylar fossa osteotomy that disrupts the TMJ, whilst trigeminal schwannoma resection approaches do not impact the TMJ. A retrospective chart review of all patients with skull base chondrosarcoma and trigeminal schwannomas will be first conducted using existing patient data from charts available on Plexia EMR and VGH hospital records/PCIS. Inclusion and exclusion criteria outlined above will be applied to select for retrospective chart analysis. Relevant surgical and clinical parameters that will be collected and reported on will correspond to pre-, peri- and postoperative records (see

attached data collection sheet). To assess quality of life, the SF-36 and DC/TMD Axis I and Axis II screening protocol will be administered to patients.

# 7. A case-control study: comparing long-term quality of life outcomes across different treatment modalities of surgery, radiation, and active surveillance in acoustic neuroma patients:

PI: Dr. Ryojo Akagami; Co-I's: Seika Taniguchi

Study period	Anticipated Number of Patients Reviewed	Approval	Status	Abstract/Paper/ Manuscript	Funding
July 2023 - Dec 2024	100	Yes	Active	N/A	N/A

#### Purpose

There is a lack of longitudinal health-related quality of life (HRQoL) outcomes of AN patients between the main three treatment modalities of surgery, stereotactic surgery and active surveillance that are matched for age, gender, and tumor characteristics. This study aims to fill this gap in the literature and investigate whether there are differences in the longitudinal HRQoL outcomes amongst AN patients between treatment modalities whilst limiting the above potential confounding variables. Findings from this study may guide clinicians in the management and integration of a biopsychosocial approach to patient care.

# Objectives

- To evaluate the longitudinal surgical and HRQoL outcomes of AN patients between the 3 primary treatment modalities of surgery, stereotactic radiosurgery and active surveillance.
- To compare and contrast the impact of surgery, radiation and active watching in the management of AN on the longitudinal HRQoL of patients.

# **Research Design**

A retrospective chart review of all AN patients will be first conducted using existing patient data from charts available on Plexia EMR and VGH hospital records/PCIS. Patients who have undergone radiotherapy management and meet the inclusion/exclusion criteria will first be identified where by patient demographics, tumor characteristics, clinical symptoms and SF-36

scores will be collected from the charts. Using this, a 1:1 match that satisfies the matched case control criteria will be applied for the remaining surgical and active surveillance group. Once patients are matched for the various variables, the same data obtained for the radiotherapy group will be collected. The control group will be the age matched Canadian health population SF-36 scores that have been already published.

### 8. Regression of Acoustic Schwannomas: a Predictive Model PI: Dr. Ryojo Akagami, Co-I's: Dr. Brian Westerber, Dr. Celine Hounjet

Study period	Anticipated number of patients reviewed	Approval	Status	Abstract/Paper /Manuscript	Funding
Oct 2023- Jan 2025	50	Yes	Data collection	N/A	N/A

#### Purpose

The purpose of this study is to attempt to identify factors that might contribute to spontaneous shrinkage of vestibular schwannomas.

#### Justification

Vestibular schwannoma are benign nerve sheath tumours that arise in the internal auditory canal and can extend into the skull, where they can cause worsening symptoms. They typically demonstrate slow growth over time and it is reasonable to manage many patients with observation alone. A portion of tumours under observation stay the same size, and some of these tumours will even shrink in size.

#### **Research Design**

A retrospective chart review of all vestibular schwannoma patients will be first conducted using existing patient data from charts available on Plexia EMR and VGH hospital records/PCIS. The research team will access medical records for the sole purpose of collecting information including patient medical history, medications, procedures, and investigations as they relate to the vestibular schwannoma tumour. The patients will also receive mailed out forms and asked to complete a survey that outlines any other existing comorbidities, lifestyle habits, and clinical presentation to gain valuable information on factors that could contribute to the spontaneous shrinkage of the vestibular schwannoma tumour.

### 9. Milrinone for the Perioperative Management of Cerebral Hypoperfusion Complicating Moyamoya Syndrome and Moyamoya Disease

PI: Dr. Peter Gooderham, Co-I: Christy Richards (Year 2 Medical Student)

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Jun 2023- Aug 2023	Approved	200	Data collection	N/A	N/A

#### Purpose

To investigate the efficacy of milrinone in alleviating hypoperfusion complications of Moyamoya disease

#### Hypothesis

There will be significant improvement of neurological outcomes in the patient cohort that is administered milrinone intraoperatively, as compared to postoperatively.

#### Justification

Moyamoya is a rare cerebrovascular disorder that is associated in cerebrohypoperfusion, manifesting in ischemic stroke events or transient ischemic attacks, which can result in permanent neurological deficits. The efficacy of milrinone in managing complications of Moyamoya is not documented in the literature. We aim to document the degree of resolution of perioperative complications for surgical revascularization in Moyamoya performed at VGH, to better understand the efficacy of milrinone and optimal timing of administration

### **10. Right Vagal Nerve Stimulators - Efficacy and Complication for treatment of** epilepsy

**PI:** Dr. Mostafa Fatehi, **Co-I:** Dr. Farzad Moien-Arshari, Dr. Jitupam Baishya, Kenneth Ong (medical student), Ru Guo (medical student)

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
January 2000 - August 2023	Awaiting VCHRI Approval	100	Data collection	N/A	N/A

To characterize the cardiac complications and efficacy of right Vagal Nerve Stimulator implantations for treatment of epilepsy.

#### Hypothesis

There will be no significant increase in implications or decrease in efficiency in right-sided Vagal Nerve Stimulator treatment.

**Justification**There are currently no studies that compare right and left-sided Vagal Nerve Stimulator treatment for epilepsy. More cases detailing right-sided Vagal Nerve Stimulator implantation for intractable epilepsy will help better evaluate the efficacy and complications of these treatments.

# 11. Vestibular Schwannomas - Progression of Symptoms During Wait-Time for Surgical Resection

PI: Dr. Serge Makarenko, Co-I: Dr. Ryojo Akagami, Dr. Celine Hounjet, Kenneth Ong (Medical Student)

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
January 2000 - August 2023	Approved	100	Data collection	N/A	N/A

The number of patients diagnosed with vestibular schwannomas has been increasing and this condition involves several progressive symptoms, including sensorineural hearing loss. The current management of vestibular schwannomas involves surgical resection, radiation, and observation, with microsurgery being the treatment of choice for larger tumors. Because vestibular schwannomas are often slow growing and considered benign, observation followed by surgery is a common approach. These surgeries may not be considered urgent operations, and the time from surgical decision to the operation does not appear to be well delineated in the literature. There also appear to be no studies that characterize the change in symptoms while patients are on a waitlist for surgery. Our study aims to characterize the progression of vestibular schwannoma symptoms for patients on a surgical waitlist, and explore whether there is a level of clinical deterioration and condition progression during this time. This knowledge may help us better understand the ideal timeline to surgery.

### Hypothesis

We hypothesize that longer wait-times between deciding when to operate and operating will lead to a higher rate of progression of symptoms in patients with vestibular schwannoma.

#### Justification

There is limited research outlining the effect of wait-times on the progression of vestibular schwannomas in patients waiting for surgical treatment for this condition. Evaluating the progression of symptoms during this wait-time can aid in understanding the ideal time to surgery and help decrease the number of symptoms experienced by the patient prior to their surgery.

### 12. Comparing Outcomes of Patients Undergoing After-Hours and Weekend Skull Base Neurosurgery at Vancouver General Hospital

**PI:** Dr. Serge Makarenko, **Co-I:** Dr. Michael Rizzuto, Amardeep Sekhon (medical student), Karanpreet Dhillon (medical student)

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
January 2000 - August 2023	Approved	200	Data collection	N/A	N/A

Surgical complications are a common cause of preventable morbidity and mortality after medication-related complications. Such complications can lead to significant healthcare costs, and it is necessary to identify preventable complications that are amenable to intervention and correction. The 'after-hour' effect refers to the idea that after-hours (i.e on-call duty) and weekend admission of surgical patients is associated with perioperative morbidity and morality rates, which may be due to surgical team fatigue, prolonged working hours, and sleep deprivation. Various studies have explored the effect of after-hour surgical start times on outcomes in spine surgery, cerebrovascular neurosurgery, and general neurosurgery, but none have looked at outcomes in cranial neurosurgery with specific emphasis on skull-base approaches. We aim to document perioperative morbidity and mortality for patients undergoing cranial neurosurgery both in-hours and after-hours, and explore whether the surgical start times have any impact on surgical outcomes.

### Hypothesis

We hypothesize that the rate of perioperative complications, surgical duration, LOS and inhospital mortality will be higher in the group of patients undergoing after-hours cranial neurosurgery compared to the in-hours group.

#### Justification

There is limited research on the effect of after-hour skull base neurosurgery on patient outcomes. Evaluating the outcomes in patients undergoing skull base neurosurgery during the daytime versus after-hours will improve our understanding on how working hour time can affect clinical outcomes.

# 13. Risk of post-craniotomy tension pneumocephalus occurring from commercial air flight

**PI:** Dr. Serge Makarenko, **Co-I:** Dr. Stefan Lang, Dr. Alexander Rebchuk, Amardeep Sekhon (student)

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Jan 2014 - Dec 2023	Awaiting REB approval	200	Awaiting approval	N/A	N/A

It is unclear whether patients can fly safely after craniotomy treatment as there is a risk of developing "tension pneumocephalus," which is classified as a neurosurgical emergency that would prompt intervention. In theory, there is an increased risk of developing post-craniotomy tension pneumocephalus from air travel based on principles in the physics of pressure system dynamics between the intracranial space and the lower-pressure aircraft cabin, as this can lead to intracranial expansion of air. There is also conflicting evidence in the literature regarding such development, causing difficulties in estimating the actual risk of developing tension pneumocephalus associated with air travel. This retrospective chart review aims to inform future post-operative care guidelines regarding air travel.

### Hypothesis

We hypothesize that there is no significant increase in risk of developing symptomatic pneumocephalus during air travel due to current air safety regulations requiring cabins of commercial aircraft at cruising altitude to be pressurized between a minimum of 75 kPa and a maximum of 101.3 kPa. As well, we hypothesize that pressure changes between 75 and 101.3 kPa are insufficient in causing significant changes in intracranial pressure attributed to the expansion of intracranial air.

#### Justification

Many patients receiving neurosurgical treatment at VGH live outside of the lower mainland in British Columbia, resulting in reliance on aviation to travel to and from home for treatment in Vancouver. This study could have clinical implications for neurosurgical post-operative care, where rates of tension pneumocephalus related to air travel can be documented and used to determine whether patients are able to travel by air shortly after discharge.

# 14. Spheno-orbital meningioma: a retrospective review of surgical and residual tumour management outcomes

PI: Dr. Serge Makarenko, Co-I: Catherine Wang (student)

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Jan 2001 - Dec 2023	Awaiting VCHRI Approval	100	Awaiting approval	N/A	N/A

Spheno-orbital meningiomas (SOM) are intracranial tumors that arise from the sphenoid wing. Surgical and clinical management for SOM is difficult due to the complexity of the associated anatomy, resulting in high risk of regrowth for residual tumors. High-dose radiation therapy has been utilized to establish residual tumor control, but can often lead to progressive visual function deterioration. This study serves the purpose of reporting the clinical course of residual SOM tumors, with a focus on tumor control, regrowth, patient visual functions, and feasibility of adjuvant irradiation, for the optimization of SOM management.

#### Hypothesis

We hypothesize that surgical morbidity in SOMs can be correlated with the tumor's resectability and that adjuvant radiation therapy provides limited benefit in terms of clinical outcomes.

#### Justification

Meningiomas are the most common type of brain tumor, and the anatomical complexity associated with SOM limits the extent of tumor resection, leading to significant morbidity. There is a higher risk of residual tumor regrowth for SOMs, leading to visual defects and significant morbidity. This study aims to aid in the optimization of SOM management by exploring the clinical course of residual SOM tumors, with a focus on tumor control and regrowth.

# 15. Primary CNS Lymphoma: is frozen section diagnosis concordant with final diagnosis?

PI: Dr. Serge Makarenko, Co-I: Ru Guo (student)

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Jan 2001 - Dec 2023	Approved	20	Data Collection	N/A	N/A

#### Purpose

Central Nervous System lymphoma comprises two main categories: secondary CNS involvement stemming from systemic lymphoma, and primary CNS lymphoma (PCNSL), where lymphoma is

confined to the brain, leptomeninges, spinal cord, or eyes, without any initial signs of involvement outside the central nervous system. In the case of secondary CNS lymphoma (SCNSL), the diagnosis is usually made from tissue collected from a systemic lesion. Conversely, PCNSL is a rare, distinct central nervous system (CNS) malignancy with unique treatment and prognostic considerations compared to other brain tumors.

Historically, PCNSL was primarily treated with whole brain radiation therapy (WBRT), akin to gliomas and other primary brain tumors. However, treatment strategies have evolved over the years, and the current approach involves initial polychemotherapy using high-dose methotrexatebased regimens, which differ from those employed for systemic NHL treatment. A key area that remains unexplored is whether the intraoperative frozen section diagnosis aligns with the final diagnosis in these tumors. Surgical debulking is recommended for glial or metastatic neoplasms, whereas PCNSL management is limited to biopsy followed by chemotherapy. Therefore, precise frozen section diagnosis is crucial for optimizing both surgical and non-surgical interventions in patients with suspected PCNSL.

### Hypothesis

We anticipate that there will be concordance between intraoperative pathological diagnoses and the final pathological diagnoses.

# 16. What quality of end-of-life care do neurosurgical patients receive, and can we do better?

PI: Dr. Serge Makarenko, Co-I: Madeline Elder (student)

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Jan 2000 - Dec 2023	Approved	100	Data Collection	N/A	N/A

#### Purpose

Research exploring the quality of end-of-life (EOL) care has been advancing and conditions for high quality EOL care have been explored in multiple studies. Consultations with specialized palliative care teams have also been shown to improve the quality of patient death, with early palliative care consults being associated with increased family satisfaction with patient death quality in literature. However, palliative care is consulted less frequently for patients requiring neurosurgical intervention compared to those hospitalized for other critical illnesses. There is no existing literature to date that explores the quality of EOL care that patients receive in neurological intensive care units in Canada. We aim to characterize the nature and timing of discussions regarding EOL care at Vancouver General Hospital (VGH) and evaluate the quality of EOL care according to standard metrics in the literature.

### Hypothesis

We hypothesize that there will be low rates of referral to palliative care and significant variation in content and contexts of EOL conversations with neurosurgical patients

#### Justification

There is a need to understand and improve the quality of EOL care that patients requiring neurosurgical care are receiving, as the mortality rate for patients admitted to the neurological intensive care unit is relatively high, and there is no research to date that explores the quality of EOL care for patients in neurological intensive care units in Canada.

# 17. Does collaboration between neurosurgery and neurotology yield better outcomes compared to neurosurgery alone in the retrosigmoid approach to vestibular schwannoma resection?

PI: Dr. Serge Makarenko, Co-I: Dr. Ryojo Akagami, Kurbaan Shergill (student)

Study period	Approvals UBC CREB/VCHRI	Anticipated number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Jan 2001 - Dec 2023	Approved	100	Data Collection	N/A	N/A

# Purpose

Vestibular schwannomas are a type of primary benign tumor that are commonly associated with asymmetrical sensorineural hearing loss, tinnitus, and facial paresthesia. Microsurgical resection is necessary once symptoms and tumor size warrant intervention. The retrosigmoid resection is a common approach to this treatment, as it offers the possibility of hearing protection by sparing the third cranial nerve, and is the only approach that can be performed by neurosurgeons alone,

whereas other resection-based treatments such as the middle cranial fossa and translabyrinthine resections of vestribular schwannoma are performed by neurosurgical and neurotological teams in conjunction.

#### Hypothesis

We hypothesize that neurotology involvement alongside the neurosurgeon in performing retrosigmoid resection of vestibular schwannomas will have improved outcomes compared to retrosigmoid resections of vestibular schwannomas performed by neurosurgeons alone.

#### Justification

It has been reported that there is insufficient evidence that a multidisciplinary team consisting of a neurosurgeon and neurotologist provides superior outcomes compared to either subspecialist working alone in the treatment of vestibular schwannoma. There have been no studies comparing results between a team-based and individual approach.

# **4. INACTIVE OR COMPLETE STUDIES**

1. Survival and Recurrence Outcomes Following Adjuvant Radiotherapy for Grade 2 Intracranial Meningiomas: 13 year Experience in Tertiary-Care Centre

PI: Dr. Serge Makarenko

Co-I: Dr. Alex Rebchuk

Study period	Approvals UBC CREB/VCHRI	Number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Oct 2007 - Oct 2021	Approved	189	Complete	Published	N/A

# **Purpose of research**

In this study, we will explore whether adjuvant radiation therapy (RT) provides overall survival (OS) and progression-free survival (PFS) benefit following resection of grade II meningioma in our local cohort. We will perform subgroup analysis to compare whether the extent of resection and timing of RT have a modifying effect on OS and PFS in this cohort. We also hope with subgroup analysis, to clarify if specific pathologic features of atypical meningiomas affect outcomes. Furthermore, we will explore whether the 2016 diagnostic change to grade II meningioma has affected the prevalence rate of these diagnoses. These results will guide local practice patterns in patients with grade II meningiomas.

# **Research Aims**

1) Explore whether adjuvant RT following operative resection of grade II meningioma has an effect on PFS and OS

a) Explore any modifying effect by Simpson grade (Grade 1 v Grade 2-3)

b) Explore any modifying effect of RT timing (early [<6wks] v late [>6wks])

2) Determine whether grade of resection predicts OS or PFS in grade II meningiomas

3) Explore clinical and histopathological predictors of OS and PFS, and tumour recurrence in grade II meningiomas

4) Explore whether recent changes to histopatholgoical diagnosis of meningiomas has affected prevalence rate

5) If possible, explore the effect of watch and wait versus operative +/- RT treatment for asymptomatic grade II meningiomas

# Design

We will perform a single center retrospective chart review. Patient level data will be obtained from Vancouver General Hospital and BC Cancer Agency database between October 2007 and December 2020.

#### Published in World Neurosurgery May 2022 Edition. Congratulations on your publication!

# 2. Recurrent Malignant Gliomas - Is there an increased perioperative risk if undergoing adjuvant chemotherapy?

PI: Dr. Serge Makarenko, Co-I: Dr. Celine Hounjet, Kenneth Ong (student)

Study period	Approvals UBC CREB/VCHRI	Number of patients reviewed	Status	Abstract/Paper /Manuscript	Funding
Jan 2001 - Dec 2022	Approved	174	Complete	Published	N/A

#### Purpose

Currently there is no standardized treatment course for patients with recurrent malignant gliomas, and treatment is mainly palliative. As repeat surgery for malignant gliomas shows increases likelihood of perioperative complications, it is important to understand the factors that pre-dispose people to these complications, with consideration to the reduced likelihood of receiving effective adjuvant radio-chemotherapy that is associated with perioperative complications. The rates and types of perioperative complications will be documented for the purpose of investigating any differences correlated with being on adjuvant chemotherapy at the time of surgery.

#### Hypothesis

We hypothesize that the rates of peri-operative complications will increase in patients undergoing surgical resection for recurrent high-grade gliomas when they are on adjuvant chemotherapy.

#### Justification

High grade gliomas are a common brain malignancy that are often recurrent, with poor prognoses for patients. Treatment for these recurrent tumors are not standardized but typically include surgery with adjuvant radio-chemotherapy, and perioperative complications for resections are not well documented in the literature. A greater understanding of risk factors around these perioperative complications and how they differ based on involvement of adjuvant radio-chemotherapy will help inform clinical treatment and utility.

Published in Journal of Neurosurgery: Case Lessons on October 16, 2023. Congratulations on your publication!

**3.** Next Generation Sequencing for Rare Variants in Familial Intracranial Aneurysm PI: Dr. William Gibson (UBC Medicial Genetics; BCCHR), Co-I's: Drs. Redekop, Haw, Gooderham

Funding	Source	Study period	Anticipated Enrolment	# of subjects enrolled	Approvals	Abstract/P aper/Manu script
Yes	HSFC	Aug 2017 - Sep 2023	Perpetual	285	Yes	N/A

#### Purpose

Several genes that predispose to aneurysms of the large blood vessels like the aorta are already known, and there are some rare genetic syndromes that predispose to brain aneurysms when other medical features (such as kidney cysts) are also present. However, there are no genes yet known that cause non-syndromic brain aneurysms. Our goal is to identify the first human gene(s) for isolated intracranial berry aneurysms.

#### **Objectives**

Our two specific aims are to catalogue the spectrum of rare coding variants in families diagnosed with intracranial aneurysms, and to validate functional effects of the most promising variant(s) on cerebral vasculature using animal models.

Study has no longer proceeded with data collection at VGH site.

4. Vertebral Artery Mobilization During Transcondylar Extreme Far Lateral Approach for Excision of Large Foramen Magnum Meningioma: 2-Dimensional Operative Video

**PI:** Dr. Ryojo Akagami, **Co-I's**: Drs. Jeremy Kam, Mendel Castle-Kirszbaum, Celine Hounjet, Benjamin Brakel, Serge Makarenko, Peter Gooderham

#### Background

Foramen magnum meningiomas are challenging lesions owing to their proximity to the lower brainstem, vertebrobasilar system, and lower cranial nerves. Tumor size, origin, morphology, relationship to neurovascular structures, and bony anatomy determine the optimal surgical approach. Classically, far lateral approaches have been the workhorse approach to the foramen magnum. Variations of the far lateral including transcondylar and extended transcondylar (paracondylar), with or without transposition of the vertebral artery, are sometimes used for a more lateral approach to the brainstem and clivus.

This video demonstrates in detail the steps of exposure, condylar drilling, vertebral artery transposition, and dural opening. These maneuvers can be difficult to conceptualize yet are key to successful extended transcondylar exposure.

#### **Video Content**

Patient presenting with a large foramen magnum meningioma. Preoperative workup includes computed tomography and MRI with angiography to assess for posterior circulation dominance, anatomic variants including posterior inferior cerebellar artery origin, venous, and bony anatomy. An extreme far lateral provides access anterior to the vertebral artery to extend exposure beyond the standard far lateral approach. This comprised transcondylar drilling, bony mobilization of the V3 Vertebral artery from C1 foramen transversarium, and dural mobilization of vertebral artery with a dural cuff at its site of dural entry.

Published in Operative Neurosurgery March 2024. Congratulations on your publication!