Lambl’s Excrescences and Stroke: A Scoping Study

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Abstract

Stroke / Cerebrovascular accident (CVA) is a leading cause of morbidity and mortality in the world. Ischemic stroke accounts for 87% of the cases, 14-30% of which is attributed to cardio-embolic stroke. Lambl’s excrescences (LE) were first described in 1856 by a Bohemian physician-VilemDusanLambl and is considered a rare cause of cardioembolic stroke subtype. LE are branched filiform structures with undulating movements, 1x4-10 mm in size that are usually found on aortic and mitral valves. An atheroma from LE or LE fragments per se may embolize to cerebrovascular arterial territory causing stroke. Multiple isolated cases of stroke associated with LE have been reported in the literature. We hereby report a scoping study of the findings associated with such cases. A total of 27 cases were identified after various scientific databases including PubMed and Google scholar were searched with keywords “lambi’s excrescences, stroke, cerebrovascular accidents”. Data from these cases were tabulated and analyzed. The mean age at presentation was 51 ± 14.2 (± SD) years with 55% of patient younger than 55 years of age. 56% of cases were males. Transesophageal echocardiogram was more effective in detecting LE when compared to transthoracic echocardiogram. LE were most often found on aortic valve and LE related stroke was most often noted in middle cerebral artery territory. Recurrent stroke was reported in 30%. Management of these cases was highly variable and likely derived from individual experience as LE management guidelines are largely lacking. Single and dual antiplatelet therapy, anticoagulation and valvularsurgery were among the various management strategies employed. We recommend dual antiplatelet after the first episode of CVA related to LE.

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Competing Interests

The authors declare that they have no competing interests.
and an antiplatelet in combination with anticoagulation after the second CVA attributed to LE. Also it is reasonable to offer valve replacement after second CVA related to LE as the recurrence rate of CVA is high. Due to rarity in LE reporting and its management a shared decision making has to be made depending on the clinical status of the patient.

The formation of a worldwide registry for LE using standardized reporting criteria for the diagnosis with or without incident stroke, would help establish guidelines for the diagnosis and management of this rare, yet serious disease with increased risk of morbidity and mortality.

Keywords
Stroke; Neurofeedback; Auditory; Upper extremity; Motor imagery

Introduction
Cerebrovascular accident (CVA) is a known cause of significant morbidity and mortality in the United States [1]. Ischemic stroke accounts for majority of strokes [2]. Cardio-embolic strokes account for nearly 14-30% of ischemic stroke and are often associated with recurrence and high mortality [3]. Lambl’s excrescences (LE) (Figure 1) though often reported in transesophageal echocardiography of patients who suffered stroke it is not clear if they are bystanders or the cardio-embolic source of such stoke [4]. Multiple isolated case reports of stroke where LE has been attributed as the cause of stroke have been reported. Guidelines regarding management of LE in the setting of stroke are largely lacking. We here undertake the first scoping study of cases of stroke associated with LE reported to understand the clinical profile of these patients, imaging studies and management that were employed, recurrent rates and mortality.

Methods
On 28th January 2018, a systematic search was conducted using Pubmed, Google Scholar, CINAHL, Cochrane CENTRAL and Web of Science databases (Figure 2). Studies listing the keywords “lambi’s excrescences, stroke, cerebrovascular accidents” were used to identify cases of stroke associated with LE. No specific duration was chosen during search. The reference list of each report was also checked for additional cases. All cases were reviewed in detail. Data reviewed included demographic data, CV risk factors, electrocardiography (ECG) findings, transthoracic echocardiography (TTE), transesophageal echocardiography (TEE), computed tomography of the head, recurrence of stroke, management of LE and death (Figure 3).

Results
A total of 27 cases were identified (Table 1) [5- 27]. The patients were in the age group of 25 to 78 years and the mean age was 51 ± 14.2 years, median age was 53 years and 44% of the cases were younger than 50 years of age and 55.5% of cases were younger than 55 years. 56% of the cases were reported in males and 44% in females. Prevalence of cerebrovascular risk factors and co-morbidities in the population was as follows: hypertension 29.63%,
diabetes 18.52%, history of stroke 14.81%, smoking 14.81%, alcohol abuse 11.11%, chronic obstructive pulmonary disease (COPD) 7.41%, history of transient ischemic attack 3.70%, history of coronary artery disease (CAD) 3.70%, obstructive sleep apnea 3.70%, deep vein thrombosis 3.70% and cocaine use 3.70%. Majority of patients presented with dysarthria/aphasia (51.85%) and hemiparesis (37.04%); other presenting complaints have been tabulated (Table 2).

TTE was reported in 14 cases, of which 6 cases (42.85%) reported LE (2 on aortic valve, 4 on mitral valve), 1 case (7.14 %)raised the suspicion for LE and no LE was detected on TTE in 7 cases (50%) all of whom were noted to have LE on TEE. TEE was reported in 22 cases of whom LE was noted on aortic valve in 82%, on mitral valve in 14% and pulmonary valve in 4%. A total of 26 cases reported the valve of LE origin which was aortic valve 73%, mitral valve 23% and pulmonary valve 4%. 25 cases reported the number of LE of which 64% of cases reported a single LE and remaining 36% of cases reported more than one LE as visualized in TEE or TTE (Table 3). Middle cerebral artery (MCA) territory was most commonly affected (Table 1). 30% reported recurrence. Management of stroke secondary to LE (Table 1 and Table 4). No mortality was reported.

Discussion

Cerebrovascular accidents (CVA) also known as stroke is a significant cause of morbidity and mortality in the United States and is the fifth leading cause of death [1]. About 800,000 cases of stroke are reported annually of which about 600 are new strokes [1]. About 87% of CVA are ischemic in nature [28-31]. When compared to firstmultiple isolated cases of LE association with stroke and recurrent stroke have been reported. Also cases of thromboembolism into coronary artery [32], renal artery [33] and popliteal artery [34] have been reported. LE usually develop at the site of valvular closure where endothelial damage happens secondary to valve wear and tear forces [35]. LE on gross examination are filiform projections with narrow base, that are 1X 4-10 mm in size and often branches are noted. LE most commonly noted on aortic and mitral valves [18, 35]. Potion of LE may detach to cause embolisation or an atheroma originating on LE may embolize to cause stroke [18, 36]. Transesophageal echocardiogram is more specific in detection of LE as compared to transesophageal echocardiogram [18, 36, 37]. Differential diagnosis for LE include i) papillary elastomers which has short broad stalk, jelly like homogeneous masses that appear in endocardium areas of valves where there is less wear and tear, ii) infective endocarditis, iii) artifact, iv) thrombus, v) redundant leaflet of a valve, and vi) intimal flap of aortic dissection [11]. Computed tomography is helpful in differentiating LE from its differentials [18, 36]. LE was noted to be present in 22% of patients on whom transesophageal echocardiogram was performed following stroke [4]. Despite the high prevalence of LE in patients who had stroke it is unclear if LE is the etiology or an incidental finding in these patients [38].

Recent trends suggest age at relatively younger age and in 2005 age at stroke was 69.2 years with 12.9% of patients under age 55 years [39]. The mean age at stroke in our study was 51 ± 14.2 years and about 55% of patients were younger than 55 years of age. Compared to statistics from 2005 [17] our patients were relatively younger likely due to cardio-embolic...
source of stroke. MCA territory was most commonly affected territory. TTE was not effective in detecting LE and missed LE in 50% of the cases and LE in these. Thus it can be concluded that TEE is superior to TTE in diagnosing LE. LE originated from aortic valve most commonly followed by mitral and pulmonary valve. Recurrent stroke was reported in 30% of cases. Though no mortality was not reported at the time of publication of these cases it is not clear if stroke secondary to LE led to mortality in any of the patients. Management strategy employed varied widely likely based on individual experience as large.

Conclusion

LE is a rare cause of cardioembolic stroke. When noted on imaging studies it is difficult to conclude if LE is the cause of stroke or just an innocent bystander. As noted in our study LE is associated with high risk of stroke recurrence. TEE is superior to TTE in detection of LE. The study design likely attributes to no mortality not noted in our study and as noted with other causes of cardioembolic stroke, LE may be expected to carry a high mortality rate on follow up. We recommend dual antiplatelet after the first episode of CVA related to LE and an antiplatelet in combination with anticoagulation after the second CVA attributed to LE. Also it is reasonable to offer valve replacement after second CVA related to LE as the recurrence rate of CVA is high. Due to rarity in LE reporting and its management a shared decision making has to be made depending on the clinical status of the patient.

Acknowledgments

Funding

This work is supported, in part, by the efforts of Dr. Moro O. Salifu M.D., M.P.H., M.B.A., M.A.C.P., Professor and Chairman of Medicine through NIH Grant number 921MD012474.

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Figure 1:
Lambl's Excrescences (indicated by green arrow) noted on aortic valve as noted in transesophageal echocardiography.
Figure 2:
Flow chart summarizing the method of selection of cases for the study.

Key words used for search: Lambl’s excrences, stroke, cerebrovascular accidents
Search conducted in: PubMed, Google Scholar, CINAHL, Cochrane CENTRAL and Web of Science databases

A total of 162 papers appeared on the search

27 papers in English literature which reported cases of LE related CVA were chosen for the scoping study
Figure 3:
Recommendations for the management of LE.
Cases of LE included in the study.

|   | Aspirin | Second antiplatelet | Statins | Anticoagulation | Valvular surgery |
|---|---------|---------------------|---------|----------------|------------------|
| 1. | 1997, Nighoghossian et al. [5] | LMCA | ✓ | ✓ | ✓, after recurrent stroke |
| 2. | 1997, Nighoghossian et al. [5] | LMCA | ✓ | ✓ | ✓, after recurrent stroke |
| 3. | 1997, Nighoghossian et al. [5] | LMCA | ✓ | ✓ | ✓, after recurrent stroke |
| 4. | 1999, Vorgos et al. [6] | RMCA | ✓ | ✓ | ✓ |
| 5. | 2003, Aggarwal et al. [7] | RMCA | ✓ | ✓ | ✓ |
| 6. | 2006, Siles Rubio et al. [8] | LMCA | ✓ | ✓ | ✓ |
| 7. | 2007, Aziz et al. [9] | LMCA | ✓ | ✓ | ✓ |
| 8. | 2007, Wolf et al. [10] | LMCA | ✓ | ✓ | ✓ |
| 9. | 2010, Kalavakunta et al. [11] | Bilateral B/L cerebral hemispheres | ✓ | ✓ | ✓ |
| 10. | 2011, Raju et al. [12] | Right posterior circulation | ✓ | ✓ | ✓ |
| 11. | 2011, Manolakis et al. [13] | Right frontal and right cerebellar | ✓ | ✓ | ✓ |
| 12. | 2012, Liu et al. [14] | Left cerebellar hemisphere | ✓ | ✓ | ✓ |
| 13. | 2012, Liu et al. [14] | Corpus Callosum, right Temporal-Parietal lobe | ✓ | ✓ | ✓ |
| 14. | 2013, Al-Amsari et al. [15] | Right Frontoparietal, left Thalamus | ✓ | ✓ | ✓ |
| 15. | 2013, Wu et al. [16] | Left posterior cerebral artery | ✓ | ✓ | ✓ |
| 16. | 2014, Veloria et al. [17] | Left central pons and corpus callosum | ✓ | ✓ | ✓ |
| 17. | 2014, Yacoub et al. [18] | LMCA | ✓ | ✓ | ✓ |
| 18. | 2015, Kitchool et al. [19] | Right thalamus, posterior capsule | ✓ | ✓ | ✓ |
| 19. | 2015, Chu et al. [20] | Right Corona radiata, Centrum Semiovale, right cerebral hemisphere | ✓ | ✓ | ✓ |
| 20. | 2015, Villlela et al. [21] | Left optic artery | ✓ | ✓ | ✓ |
| 21. | 2015, Dumitrascu et al. [22] | Left parietal lobe, bilateral centrum semiovale, right occipital | ✓ | ✓ | ✓ |
| 22. | 2015, Davoguesto et al. [23] | Left centrum Semiovale | ✓ | ✓ | ✓ |
| 23. | 2016, Kamran et al. [24] | LMCA | ✓ | ✓ | ✓ |
| 24. | 2016, Jo et al. [25] | RMCA | ✓ | ✓ | ✓ |
| 25. | 2016, Jo et al. [25] | LMCA | ✓ | ✓ | ✓ |
| 26. | 2017, Meireles et al. [26] | Right occipital Lobe, cerebellar hemispheres | ✓ | ✓ | ✓ |
|   | Aspirin | Second antiplatelet | Statins | Anticoagulation | Valvular surgery |
|---|---------|---------------------|---------|-----------------|-----------------|
| 27. | 2017, Vlacancich et al. [27] | Right precentral gyrus | ✓       | ✓               |                 |

LMCA= left middle cerebral artery.
RMCA= right middle cerebral artery
Table 2:

Presenting complaint (n=27).

| Complaint                      | Percentage |
|--------------------------------|------------|
| Dysarthria/aphasia             | 51.85%     |
| Hemiparesis/paresis            | 44.45%     |
| Headache                       | 18.52%     |
| Altered mental status          | 14.81%     |
| Dizziness                      | 11.11%     |
| Facial droop                   | 7.41%      |
| Blurry vision                  | 7.41%      |
| Photophobia                    | 7.41%      |
| Phonophobia                    | 7.41%      |
| AmaurosisFugax                 | 3.70%      |
| Falls                          | 3.70%      |
| Diplopia                       | 3.70%      |
Table 3:

| Serial number | Year, author                          | Transthoracic echocardiogram | Transesophageal echocardiogram |
|---------------|--------------------------------------|------------------------------|--------------------------------|
| 1.            | 1997, Nighoghossian et al. [5]        | Normal                       | LE on mitral valve             |
| 2.            | 1997, Nighoghossian et al. [5]        | LE on mitral valve           | -                              |
| 3.            | 1997, Nighoghossian et al. [5]        | LE on mitral valve           | -                              |
| 4.            | 1999, Vorgos et al. [6]              | -                            | 4 LE on Pulmonary valves       |
| 5.            | 2003, Aggarwal et al. [7]            | LE on aortic valve           | -                              |
| 6.            | 2006, Siles Rubio et al. [8]         | -                            | LE on Aortic Valve             |
| 7.            | 2007, Aziz et al. [9]                | -                            | LE on Aortic valve             |
| 8.            | 2007, Wolf et al. [10]               | -                            | LE on Aortic Valve             |
| 9.            | 2010, Kalavakunta et al. [11]        | -                            | LE on Aortic valve             |
| 10.           | 2011, Raju et al. [12]               | Suspicious of Aortic Valve mass | LE on Aortic Valve        |
| 11.           | 2011, Manolakis et al. [13]          | -                            | LE on Aortic Valve             |
| 12.           | 2012, Liu et al. [14]                | -                            | LE on Aortic Valve             |
| 13.           | 2012, Liu et al. [14]                | -                            | LE on Aortic Valve             |
| 14.           | 2013, Al-Ansari et al. [15]          | LE on Mitral Valve           | LE on Mitral Valve             |
| 15.           | 2013, Wu et al. [16]                 | -                            | LE on Aortic Valve             |
| 16.           | 2014, Veloria et al. [17]            | -                            | LE on Aortic Valve             |
| 17.           | 2014, Yacoub et al. [18]             | Normal                       | LE on Aortic Valve             |
| 18.           | 2015, Kitchool et al. [19]           | Normal                       | LE on Mitral Valve             |
| 19.           | 2015, Chu et al. [20]                | Normal                       | LE on Aortic Valve             |
| 20.           | 2015, Villella et al. [21]           | LE on Mitral Valve           | -                              |
| 21.           | 2015, Dumitrascu et al. [22]         | Normal                       | LE on Aortic Valve             |
| 22.           | 2015, Davogusto et al. [23]          | LE on Aortic Valve           | LE on Aortic Valve             |
| 23.           | 2016, Kantran et al. [24]            | Normal                       | LE on Aortic Valve             |
| 24.           | 2016, Jo et al. [25]                 | -                            | LE on Aortic valve             |
| 25.           | 2016, Jo et al. [25]                 | -                            | LE on Aortic valve             |
| 26.           | 2017, Meireles et al. [26]           | -                            | -                              |
| 27.           | 2017, Vlacancich et al. [27]         | -                            | LE on Aortic valve             |
Table 4: Management of stroke secondary to LE.

| Treatment                        | Percentage |
|---------------------------------|------------|
| Aspirin                         | 30%        |
| Dual antiplatelet therapy       | 19%        |
| Anticoagulation                 | 30%        |
| Valve surgery                   | 30%        |