Non Rheumatic Atrial Fibrillation as Risk of Stroke

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To cite this article:
Ali Areef Fadhlullah, Asgad A. Abdalgbar, Hanan K. Altalhi. Non Rheumatic Atrial Fibrillation as Risk of Stroke. American Journal of Internal Medicine. Vol. 4, No. 6, 2016, pp. 117-119. doi: 10.11648/j.ajim.20160406.15

Received: October 21, 2016; Accepted: November 1, 2016; Published: November 25, 2016

Abstract: Atrial fibrillation (AF) is one of the main risk factor for ischemic stroke. The reason for an increased stroke risk in AF has always been claimed to be the occurrence of left atrial thrombosis causing arterial embolism. In patients with Rheumatic heart disease especially mitral valve stenosis with AF, the frequency of atrial thrombosis has found to be 30 - 42% (Keren G et al. 1987), and the prevalence of left atrial thrombosis in NRAF are higher than in control 13-27% (Petersen P et al. 1988). Objectives: We investigated if there are any differences in risk factors or atherosclerotic manifestations between ischemic brain infarction patients with and without AF? Are the brain lesions characteristics in these patients similar? Patients and Methods: This is a case - control study of 26 patients with acute ischemic stroke and NRAF (case subjects) a and 26 patients with acute ischemic stroke and sinus rhythm. (control subjects). The patients admitted to the hospital; the diagnosis of cerebral infarction was confirmed by new CT of the brain. All the participants underwent the standard examination and testing as well as ECG and ECHO. Result: Patient with NRAF had higher mortality 30% than in control (SR) 7% (P<0.001). NRAF patients had positive brain CT finding 68% compared to 23% of the SR Patients (P<0.001). Conclusion: Brain infarction in non-Rheumatic AF group are more serious than other and therefor make up a (high risk) group for which acute treatment would be specially indicated and primary prophylaxis with anticoagulants might therefore be considered.

Keywords: Cerebral Vascular Accident (CVA), Non Rheumatic Atrial Fibrillation (NRAF), Sinus Rhythm (SR), Electrocardiography (ECG), Echocardiography (ECHO), Atrial Fibrillation (AF)

1. Introduction

Atrial fibrillation has been shown to be associated with increased risk for cardiovascular disease morbidity (Britton M, et al. 1985) and mortality (Kannel WB, Et al. 1982). Although the importance of AF partly derives from its strong association with ischemic stroke. There have not been as many advances in our understanding of the mechanism of stroke in AF. Current views rest on a century old hypothesis that fibrillation of the atrium produce stasis of blood which cause thrombosis formation and embolism to the brain. When other abnormalities are acknowledged to play role, the dysrhythmia is still considered the primary cause of, thromboembolism (Zimetbaum P, et al. 2014). Non rheumatic atrial fibrillation carries a nearly fivefold increased risk for stroke compared with normal sinus rhythm. (Wolf PA, et al. 1991).

2. Objectives

We investigated if there are any differences in Risk factors or atherosclerotic manifestations between brain infarction patients with and without AF?

Are the brain lesions characteristics in these patients similar?

3. Patients and Methods

This is case - control study of patients admitted to the hospital with diagnosis of acute ischemic infarction. Data were collected with interview the cases, physical and neurological examination done by study physicians. The diagnosis of ischemic stroke was confirmed by new computer Tomography (CT).

ECG done for every CVA patients as interpreted by
cardiologist from standard 12 leads ECG, the patients were divided to two groups according to the ECG finding, chronic AF (n=26), constant AF during hospital stay, sinus rhythm (n=26)-not known episodes of AF during hospital stay.

The ECG was appraised according to the Minnesota code (The Scandinavian committee on ECG classification).

Echo: Transthoracic echocardiography (TTE) was performed in all subjects and assessed by cardiologist. 2D, M – Mode – PW - CW were performed to asses LVEF, was measured by simplified cylinder - hemiellipsoid formula (Weyman AE, et al. 1994) and excluded the Rheumatic heart Valve disease.

Other various Risk factors:
BP measurement in right arm, Cigarette smoking to be qualified based on daily consumption and duration of smoking, blood biochemistry analysis were made ie. Fasting blood sugar (FBS) total cholesterol, TG, CBC.

Statistical analysis:
Statistical analysis were performed by using SPSS software (version 11) difference among groups were analyzed by one way. unstack ANOVA and P values 0.05 were considered significant.

4. Result

Table 1: shows Characteristics and variation of study subjects: 52 patients with acute ischemic stroke were included in the study, 26 patient with NRAF (case subject ) and 26 patient with SR (control subjects). There is significant difference in mean age between NRAF group and SR group being older in NRAF group (p 0.001), and number of patients with history of diabetes and heart failure is significantly increased among NRAF group (p 0.007, p 0.001 respectively).

| Table 1. Study population characteristics and variation. |
|----------------------------------------------------------|
|                | AF | SR | P value |
| Number         | 26 | 26 |         |
| Age mean ± SD  | 68.96 ± 15.46 | 52.96 ± 16.39 | 0.001 |
| Rang           | 40 - 90 | 30 - 75 |         |
| Male           | 14 (53.8%) | 17 (65.4%) | 0.406 (NA) |
| Female         | 12 (46.2%) | 9 (34.6%) | 0.406(NA) |
| Diabetic       | 16 (61.5%) | 7 (27%) | 0.007 |
| Hypertension   | 8 (30.8%) | 4 (15.4%) | 0.162(NA) |
| Smoking        | 9 (34.6%) | 11 (42.3%) | 0.724 (NA) |

| Table 2. Clinical, Laboratory, ECHO and ECG data of study groups. |
|---------------------------------------------------------------|
| Initial SBP mmHg | Initial DBP mmHg | ECG % |
| AF n=26          | SR n=26          |      |
| 149.15 ± 36.65   | 148.08 ± 37.31   | NA    |
| 92.69 ± 25.07    | 88.27 ± 16.31    | NA    |
| 7               | 16                | P<0.05 |
| Sign MI          |                  |      |
| 8               | 5                 | NA    |
| LVH             |                  |      |
| 3               | 4                 | NA    |
| LBBB            |                  |      |
| 5               | 2                 | NA    |
| Normal (except rhythm) |          |      |
| 16 (61%)        | 23 (88%)         | NA    |
| Triglycerides   |                  |      |
| 91.33 ± 25.88   | 102.14 ± 42.12   | NA    |
| Cholesterol     |                  |      |
| 150 ± 61.69     | 127.20 ± 76.98   | NA    |
| 91.33 ± 25.88   | 102.14 ± 42.12   | NA    |
| 5               | 2                 | NA    |
| 10 (25%)        | 3 (7.5%)         | P 0.0001 |
| FBS             |                  |      |
| 150 ± 61.69     | 127.20 ± 76.98   | NA    |
| 91.33 ± 25.88   | 102.14 ± 42.12   | NA    |
| NA             |                  |      |
| 150 ± 61.69     | 127.20 ± 76.98   | NA    |
| 91.33 ± 25.88   | 102.14 ± 42.12   | NA    |
| 5               | 2                 | NA    |
| 10 (25%)        | 3 (7.5%)         | P 0.0001 |
| LV dysfunction  |                  |      |
| 2               | 4                 | NA    |
| Cholesterol     |                  |      |
| 91.33 ± 25.88   | 102.14 ± 42.12   | NA    |
| Triglycerides   |                  |      |
| 91.33 ± 25.88   | 102.14 ± 42.12   | NA    |
| 5               | 2                 | NA    |
| 10 (25%)        | 3 (7.5%)         | P 0.0001 |
| Normal (except rhythm) |          |      |
| 16 (61%)        | 23 (88%)         | NA    |
| Triglycerides   |                  |      |
| 91.33 ± 25.88   | 102.14 ± 42.12   | NA    |
| Cholesterol     |                  |      |
| 150 ± 61.69     | 127.20 ± 76.98   | NA    |
| 91.33 ± 25.88   | 102.14 ± 42.12   | NA    |
| 5               | 2                 | NA    |
| 10 (25%)        | 3 (7.5%)         | P 0.0001 |
| LVH             |                  |      |
| 150 ± 61.69     | 127.20 ± 76.98   | NA    |
| 91.33 ± 25.88   | 102.14 ± 42.12   | NA    |
| 5               | 2                 | NA    |
| 10 (25%)        | 3 (7.5%)         | P 0.0001 |

Table 3: Shows Brain lesions characteristics. And condition on admission: In brain, infarction patients with and without AF the symptoms were known to have occurred suddenly in 60% of cases in groups. Paresis was the dominating symptoms. Seizures in the first hour occurred in (11%) of the AF group compared to (4%) of SR group (P>0.05), impaired level of consciousness occurred in (31%) of the AF group compared to (4%) of the SR group (p<0.01), patients cannot walk independently occurred in (69%) AF group compared to (31%) of the SR group (p<0.05), at admission to hospital the AF group had several signs of more serious brain damage than the SR group.

Mortality was higher among the AF group (30% vs. 23% respectively; P.<0.001).

No difference was seen between the groups as regards the location of the brain lesion and there were more pathological finding at CT brain in the AF group (68%) compared to (23%) of SR group (P =0.001).

| Table 3. Brain Lesion Characteristics in 52 Patients with NRAF and with SR. |
|---------------------------------------------------------------|
| Condition on admission | AF n=26 | SR n=26 | P value |
| Seizures in the first hours | 3 (11%) | 1 (4%) | P<0.05 |
| Impaired consciousness | 8 (31%) | 1 (4%) | P<0.01 |
| Cannot walk independently | 18 (69%) | 8 (31%) | P<0.05 |
| Mortality | 8 (30%) | 2 (7%) | P<0.001 |
| Brain CT (Positive) | 18 (68%) | 6 (23%) | P<0.001 |

5. Discussion

The AF patients were older and had more serious brain damage than those with SR. Some factors have to be dealt with before discussing whether these and other difference between the groups favors an Athero-thrombotic or embolic origin of stroke in AF patients. We chose to compare a patients with AF with those with SR. It has been claimed that a sudden onset of maximum symptoms in patients with an embolic sources, like AF is typical embolism (Easton JD, et al. 1980; Dalal PM, et al. 1965).

We found progression of symptoms after arrival at hospital to be as common among NRAF patients as among others (roden A et al. 1982).
AF has been consistently associated with stroke in different cohorts (Wolf PA et al. 1978).

AF frequently co-exists with atrial abnormalities, such as endothelial dysfunction (Cai H, et al. 2002), impaired myocyte function (Frustaci A, ey al. 1997), and mechanical dysfunction of left atrial appendage (Mihm MJ, et al. 2001), these abnormality have been documented in both experimental animal models and in human (Warraich HJ, et al. 2014), could these abnormality also arise independently of AF and cause stroke. Once AF develops the dysrhythmia cause contractile dysfunction and stasis, which further increase the risk of thromboembolism. In addition over time the dysrhythmia cause structural remodeling of the atrium there by worsening atrial cardiopathy and the risk of thromboembolic. There is strong association between AF and embolic stroke. 10% of patients with lacunar infarction have AF (loddor J, et al. 1990).

The AF patients were moreover older, reflecting the increasing prevalence of AF with increasing age. Arteriosclerotic macro-and microangiopathy progressing with age, might instead be the cause of the stroke as well as the heart disease.

The finding in this study of more severe brain damage in AF patients may also argue in favors of embolism. A sudden interruption of blood flow caused by an embolus might give rise to large infarction than gradual occlusion with opportunity for development of collateral circulation. More sever lesions in embolic stroke have also been found at autopsy (Jorgensen L, et al. 1969). AF was noted to be associated with worse prognosis (Britton M, et al. 1985). There might be other explanation than embolism for the poor outcome of stroke in AF patients. A decreased cardiac output due to the higher incidence of heart failure might be contributed to a reduced cerebral blood flow when auto regulatory mechanism are impaired as in an ischemic region (Keller TS, et al. 1982).

### 6. Conclusion

We would like to conclude that embolism is a plausible cause of stroke in many NRAF patients. NRAF patients not only have increased incidence of stroke but their stroke are more serious for which acute treatment would be especially indicated. More work is required to determine whether additional marker such as cardiac MRI of tissue fibrosis and computed tomography assessment of left atrial appendage morphology, may better identify the risk of atrial thromboembolism

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