Aspirin for the Primary Prevention of Myocardial Infarction: An Evidence-Based Clinical Inquiry

Brooklyn R. Nemetchek*

Abstract

Evidence is evaluated to determine whether low dose aspirin (81mg) for primary prevention in patients aged 50-65 with no history of cardiovascular disease decreases the incidence of myocardial infarction. Ten studiesgiving relevant clinical evidence are identified and evaluated, with each gender looked at in isolation. The preliminary evidence of this paper suggests that aspirin for the primary prevention of myocardial infarction is not suitable for women aged 50-65, while it does hold benefits for males of the same age range (Howard, 2014). However, the evidence is not unanimous, and more research is needed before recommending aspirin for primary prevention in all low-risk individuals. In relation to aspirin for primary prevention of myocardial infarction, short- and long-term recommendations for nursing practice are developed and discussed, demonstrating the significant role the nurse plays in education, helping each patient to assess individual risks and benefits, and advising patients to consult their physician before self-medicating (Howard, 2014).

Keywords: aspirin, primary prevention, myocardial infarction, nursing

Cardiovascular disease is the number one cause of death globally, having claimed an estimated 17.5 million lives in 2012. Of those, coronary heart disease accounted for approximately 7.4 million deaths (World Organization [WHO], 2015). Cardiovascular disease describes disorders of the heart and blood vessels, including coronary heart disease, cerebrovascular peripheral arterial disease, rheumatic heart disease, congenital heart disease, and deep vein thrombosis (WHO, 2015). Myocardial infarction, a subgroup of coronary heart disease, occurs when a vessel supplying the heart is occluded, usually by a clot (WHO, 2015). A primary prevention strategy is long-term administration of aspirin for the purpose of preventing the first occurrence of cardiovascular disease, including, more particular to this

paper, myocardial infarction (Kappagoda & Amsterdam, 2011; American Society of Health-System Pharmacists [ASHP], 1997).

Aspirin (acetylsalicylic acid) is a potent and irreversible inhibitor of platelet aggregation because it reduces thrombosis (Gaziano & Greenland, 2014). In platelets, the enzyme COX-1 produces thromboxane A2 which aids in platelet aggregation (Gaziano & Greenland, 2014). COX-1 cannot be regenerated in platelets and is therefore permanently inhibited by aspirin, leading to a prolonged antithrombotic effect lasting several days after a single dose until enough new platelets have been produced to restore normal function (Gaziano & Greenland, 2014). This unique property means aspirin is valuable in reducing the risk of thrombotic events such as myocardial infarction, but

also in increasing the possibility of bleeding, including gastrointestinal bleeding and hemorrhagic stroke (Gaziano & Greenland, 2014). Aspirin has been shown to have benefits during acute events, after certain vascular procedures, and as secondary prevention of major vascular events for those with evidence of cardiovascular disease (Gaziano & Greenland, 2014). The majority of the population taking aspirin for cardiovascular prevention use it for primary prevention (Howard, 2014). Approximately 20% of these individuals do so without medical recommendation (Howard, 2014), while the question remains whether aspirin is appropriate for everyone (Howard, 2014).

Health care practitioners, and nurses in particular, have a key role in ensuring aspirin is used safely. In order to evaluate aspirin and its potential benefits and harms for each patient, the nurse must be able to identify and solve problems related to body systems, interpret physical assessments and diagnostic data, be aware of the factors affecting safe nursing practice, and communicate interprofessionally in relation to the care of complex and high acuity patients. These nursing requirements are critical to the care of such a patient. Gaziano and Greenland (2014) note that "Given the beneficial effects of aspirin during acute events, following procedures, and in the secondary prevention of major vascular events among patients with cardiovascular disease, it was logical to ask whether this inexpensive drug could prevent the first myocardial infarction or stroke among persons who have yet to manifest vascular disease". Evidence will be evaluated to determine whether or not low dose aspirin (81mg) for primary prevention decreases the incidence of myocardial infarction in patients between the ages of 50 and 65 with no history of cardiovascular disease.

Methodology

Randomized control trials (RCTs) have been conducted on the issue with varying populations, dosages of aspirin, controls, and outcomes. Meta-analysis of RCTs themselves have resulted in differences in terms of recommendations for practice. Results are based on studies of primarily white males or health care providers, which limited the generalizability of the findings (Howard, 2014). Therefore, there is a need for further study regarding particular ages, populations, and subpopulations (for example, women age 65 and older as compared to women aged 50-65). A literature review was conducted using Medline and Cumulative Index to Nursing and Allied Health (CINAHL), using the key phrases of "myocardial infarction," "primary prevention," and "aspirin." Studies looking at aspirin's effect on primary occurrence of myocardial infarction were included. Studies looking only at other potential outcomes of aspirin use, such as incidence of stroke and bleeding, were not included.

Results

Eleven studies giving relevant clinical evidence were identified and are presented in Table 3, evaluated alphabetically by author. The level of evidence for each study was recognized with the use of Table 2. To examine the evidence, it is helpful to look at each gender in isolation, as significant differences have been observed.

Table 1: PICO Question Format for Aspirin as a Prevention of Myocardial Infarction

Patient or problem	Individuals (both men and women) between the ages of 50 and 65 with no history of cardiovascular disease.
Intervention	Low-dose aspirin (81mg) for primary prevention. 81mg is the lowest readily available dose in chewable tablets and enteric-coated tablet forms (Lippincott Williams & Wilkins, 2005, p. 355). 75-100mg has been shown to be the optimum dosage range due to its maximal inhibition of platelet aggregation while giving minimal side effects (Hennekens, Manson & Reilly, 2002).
Comparison	No aspirin treatment or placebo. Comparisons made to other antiplatelet therapies, although they may be present in the research articles examined, are not included in this analysis.
Outcome	Decreased incidence of myocardial infarction. Although other outcomes such as bleeding are possible, the paper looks solely at the incidence of myocardial infarction for an indication of an outcome.

Table 2: Levels of Evidence for Prevention Medicine

Level	Type of Evidence
1a	Systematic review of Randomized Control Trials (RCTs)
1b	Individual RCT (with narrow confidence intervals)
10	All or none study
28	Systematic review (with homogeneity) of cohort studies
2b	Individual cohort study (including low quality RCT, e.g.,<80% follow-up)
2C	"Outcomes" research; Ecological studies
3a	Systematic review (with homogeneity) of case-control studies
3b	Individual case-control study
4	Case series (and poor quality cohort and case-control study)
5	Expert opinion without explicit critical appraisal, or based on physiology bench research or "first principles"

(Howick, 2009)

Table 3a: Evaluation of Source and Level of Evidence

	Article 1
Reference	Antithrombotic Trialists' Collaboration. (2002). Collaborative meta-analysis of randomized trials of antiplatelet therapy for prevention of death, myocardial infarction, and stroke in high-risk patients. British Medical Journal, 324(7329), 71-86. doi: 10.1136/bmj.324.7329.71
Level of Evidence	1a
Research Design	Collaborative meta-analyses of 287 studies (197 comparing anti-platelet verses control; 90 comparing different antiplatelet regimens).
Problem/Purpose	To determine the effects of antiplatelet therapy among patients at high risk of occlusive vascular events.
Setting	Varied.
Sample/Population	(N=135 000) patients in comparisons of antiplatelet therapy versus control. (N=77 000) patients in comparisons of different antiplatelet regimens.
Aspirin	Varied dosages.
Comparison	Placebo, no aspirin, or other antiplatelet therapy.
Findings	Aspirin (or another oral antiplatelet drug) is protective in most types of patients at increased risk of occlusive vascular events, including those with an acute myocardial infarction or ischaemic stroke, unstable or stable angina, previous myocardial infarction, stroke or cerebral ischaemia, peripheral arterial disease, or atrial fibrillation. Low dose aspirin (75-150 mg daily) is an effective antiplatelet regimen for long-term use.
Recommendations and/or Limitations	An unanswered question is whether it is possible to identify particular groups of apparently healthy people who may be at increased risk of myocardial infarction or stroke and for whom the benefits of daily aspirin outweigh the risks. Adding a second antiplatelet drug to aspirin may produce additional benefits in some clinical
	circumstances, but more research into this strategy is needed. Aspirin dose, duration of treatment, and lengths of follow-up were not uniform. Meta-analysis remains a retrospective research subject to the methodological deficiencies of the included studies.

Table 3b: Evaluation of Source and Level of Evidence

	Article 2
Reference	Bartolucci, A., Tendera, M., & Howard, G. (2011). Meta-analysis of multiple primary prevention trials of cardiovascular events using aspirin. <i>The American Journal of Cardiology</i> , 107(12), 1796-1801. doi:10.1016/j.amjcard.2011.02.325
Level of Evidence	18
Research Design	Analysis of nine randomized trials evaluating the benefits of aspirin for primary prevention of cardiovascular events: - The British Doctor's Trial (BMD) - The Physician's Health Study (PHS) - The Thrombosis Prevention Trial (TPT) - The Hypertension Optimal Treatment (HOT) study - The Primary Prevention Project (PPP) - The Women's Health Study (WHS) - The Aspirin for Asymptomatic Atherosclerosis Trial (AAAT) - The Prevention of Progression of Arterial Disease and Diabetes (POPADAD) trial - The Japanese Primary Prevention of Atherosclerosis with Aspirin for Diabetes (JPAD) trial
Problem/Purpose	To determine the role of aspirin in primary prevention of cardiovascular events.
Setting	Varied.
Sample/Population	Both genders, varying age and medical history.
Aspirin	(N=50 868). Varying dosage: 75-325mg
Comparison	(N= 49 170). Placebo or no aspirin.
Findings	Primary prevention with aspirin decreased the risk for total cardiovascular events and nonfatal MI, but there were no significant differences in the incidences of stroke, cardiovascular mortality, all-cause mortality, and total coronary heart disease.
Recommendations and/or Limitations	Aspirin dose, duration of treatment, and lengths of follow-up were not uniform. Meta-analysis remains a retrospective research subject to the methodological deficiencies of the included studies.

Table 3c: Evaluation of Source and Level of Evidence

	Article 3
Reference	Berger, J., Roncaglioni, M., Avanzini, F., Pangrazzi, I., Tognoni, G. & Brown, D. (2006). Aspirin for the primary prevention of cardiovascular events in women and men: A sex-specific meta-analysis of randomized controlled trials. <i>The Journal of the American Medical Association</i> , 295(3), 306-13. doi:10.1001/jama.295.3.306
Level of Evidence	18
Research Design	Meta-analysis of six randomized controlled trials of aspirin therapy in participants without cardiovascular disease that reported data on myocardial infarction (MI), stroke, and cardiovascular mortality. - Physician's Health Study (1989) - British Doctor's Trial (1988) - Thrombosis Prevention Trial (1998) - Hypertension Optimal Treatment Trial (1998) - Primary Prevention Project (2001) - Women's Health Study (2005)
Problem/Purpose	To determine the effect of aspirin in the primary prevention of cardiovascular disease in women and men independently.
Setting	Varied.
Sample/Population	(N=95 456 total): 3 trials included only men, 1 included only women, and 2 included both sexes. 51 342 were women.
Aspirin	75mg/day (2 trials) 100mg every other day (1 trial) 100mg/day (1 trial) 325mg every other day (1 trial) 500mg/day (1 trial)
Comparison	Varied. Placebo or no aspirin.
Findings	For women and men, aspirin therapy reduced the risk of a composite of cardiovascular events due to its effect on reducing the risk of ischaemic stroke in women and MI in men. Aspirin significantly increased the risk of bleeding to a similar degree among women and men.
Recommendations and/or Limitations	Aspirin dose, duration of treatment, and lengths of follow-up were not uniform.
	Meta-analysis remains a retrospective research subject to the methodological deficiencies of the included studies.

Table 3d: Evaluation of Source and Level of Evidence

	Article 4
Reference	Hayden, M. (2002). Aspirin for the primary prevention of cardiovascular events: A summary of the evidence for the U.S. preventive services task force. <i>Annals of Internal Medicine</i> , 136(2),161-72. doi:10.7326/0003-4819-136-2-200201150-00016
Level of Evidence	28
Research Design	RCTs at least 1 year in duration examining aspirin primary prevention in patients without previously known cardiovascular disease and systematic reviews, recent trials, and observational studies.
Problem/Purpose	The use of aspirin to prevent cardiovascular disease events in patients without a history of cardiovascular disease is controversial. To examine the benefits and harms of aspirin primary prevention.
Setting	United Kingdom, United States, Italy, and Worldwide.
Sample/Population	(N=53 035) total men and women from 5studies, age range of 40-84. Included male physicians, men at high risk for heart disease, men and women with diastolic blood pressure of 100-115mmHg, and those with more than one major risk factor for coronary heart disease. (N=11 414) women in 20f the 5studies.
Aspirin	500 mg/day 325 mg every other day 75 mg/day (controlled release) 75 mg/day 100 mg/day
Comparison	Placebo or no placebo.
Findings	The net benefit of aspirin increases with increasing cardiovascular risk. Aspirin reduced the relative risk for myocardial infarction for patients 70 to 84 years of age (relative risk, 0.49) as much as or more than it did for patients 60 to 69 years of age (relative risk, 0.46) and patients 50 to 59 years of age (relative risk, 0.58).
Recommendations and/or Limitations	Most of the participants in the 5 randomized trials were middle-aged men. There is a need to examine whether the effect of aspirin differs in other demographic groups, including elderly persons, women, and persons with diabetes or hypertension.

Table 3e: Evaluation of Source and Level of Evidence

	Article 5
Reference	Howard, P. A. (2014). Aspirin for primary cardiovascular prevention: When is it worth the risks?
	Hospital Pharmacy, 49(6), 502-507. doi:10.1310/hpj4906-502
Level of Evidence	5
Research Design	Review of evidence, including trials and meta-analysis, recommendations, issues to be resolved, and
	pathophysiology.
Problem/Purpose	It has become increasingly clear that aspirin may not be appropriate for everyone. This article reviews
	current evidence, recommendations, and issues that have yet to be resolved.
Setting	Varied.
Sample/Population	Varied.
Aspirin	Varied dosage.
Comparison	Varied, including placebo.
Findings	Current evidence has not provided definitive proof of aspirin's effectiveness for primary prevention
	among those with low baseline risk.
Recommendations	Many of the clinical trials were conducted prior to the widespread use of other preventive therapies
and/or Limitations	such as statins; aspirin's effects onpatients on these drugs require further study.

Table 3f: Evaluation of Source and Level of Evidence

	Article 6
Reference	Ikeda, Y., Shimada, K., Teramoto, T., Uchiyama, S., Yamazaki, T., Oikawa, S., & Ishizuka, N.
	(2014). Low-dose aspirin for primary prevention of cardiovascular events in Japanese patients 60
	years or older with atherosclerotic risk factors: A randomized clinical trial. JAMA: Journal of the
	American Medical Association, 312(23), 2510-2520. doi:10.1001/jama.2014.15690.
Level of Evidence	2b
Research Design	Open-label, randomized (1:1), parallel-group trial over 5 years.
Problem/Purpose	To determine whether daily, low-dose aspirin reduces the incidence of cardiovascular events in older
	Japanese patients with atherosclerotic risk factors.
Setting	1007 clinics in Japan.
Sample/Population	Japanese patients (N=14464) aged 60 to 85 years, presenting with hypertension, dyslipidemia, or
	diabetes mellitus.
Aspirin	Enteric-coated aspirin, 100mg/day.
Comparison	No aspirin.
Findings	Once-daily, low-dose aspirin did not significantly reduce the risk of the composite outcome of
	cardiovascular death, nonfatal stroke, and nonfatal myocardial infarction among Japanese patients
	6o years or older with atherosclerotic risk factors.
Recommendations	Study terminated early based on a futility assessment; however, an exploratory analysis suggested a
and/or Limitations	28% probability of finding a significant difference in favour of aspirin had the study been continued
	through the planned number of events.
	There was a decreasing level of adherence with daily low-dose aspirin in the aspirin group (dropping
	to 76% in year 5) and increasing uptake of daily aspirin in the no aspirin group (10% by year 5).
	The study was not blinded and did not have a placebo control due to Japanese ethical standards, and
	, , , , , , , , , , , , , , , , , , , ,
	the drop-in rate and rate of patients lost to follow-up were about 10%. The role that enteric coamight have on the efficacy of low-dose aspirin is not clear.

Table 3g: Evaluation of Source and Level of Evidence

	Article 7
Reference	Juul-Möller, S., Edvardsson, N., Jahnmatz, B., Rosén, A., Sørensen, S., & Omblus, R. (1992). Double-blind trial of aspirin in primary prevention of myocardial infarction in patients with stable chronic
	angina pectoris. The Swedish Angina Pectoris Aspirin Trial (SAPAT) Group. <i>Lancet</i> , 340(8833), 1421-
	1425. doi:10.1016/0140-6736(92)92619-Q
Level of Evidence	1b
Research Design	Randomized, double blind.
Problem/Purpose	To compare aspirin and placebo in patients with a history of chronic stable angina pectoris without a
	previous MI.
Setting	Sweden.
Sample/Population	Patients (N=2035) aged 30-80 who demonstrated chest pain on exertion for at least 1month.
Aspirin	75mg/day and sotalol for angina symptoms.
Comparison	Placebo and sotalol for angina symptoms.
Findings	Addition of low-dose aspirin to sotalol showed significant benefit in terms of cardiovascular events,
	including a significant reduction in the incidence of first myocardial infarction in patients with
	symptoms of stable angina pectoris. 75mg aspirin in combination with sotalol decreased incidence of
	first myocardial infarction by 34%.
Recommendations	Further study is needed to determine the effects of aspirin by itself with the same population,
and/or Limitations	without the use of sotalol or other medications of its kind.

Table 3h: Evaluation of Source and Level of Evidence

	Article 8
Reference	Manson, J., Stampfer, M., Colditz, G., Willett, W., Rosner, B., Speizer, F., & Hennekens, C. (1991). A prospective study of aspirin use and primary prevention of cardiovascular disease in women. <i>The Journal of the American Medical Association</i> , 266(4), 521-527. doi:10.1001/jama.1991.03470040085027.
Level of Evidence	2b
Research Design	Prospective cohort study including 6 years of follow-up, from 1980 to 1986. Participants responded to questionnaires regarding medical history and lifestyle.
Problem/Purpose	To examine prospectively the association between regular aspirin use and the risk of a first myocardial infarction and other cardiovascular events in women.
Setting	Eleven U.S. states.
Sample/Populatio	(N = 87 678) United States registered nurses aged 34 to 65 years free of diagnosed coronary heart
n	disease, stroke, and cancer at baseline.
Aspirin	Varying amounts. Participants were asked whether or not they took aspirin, and then how many aspirin per week (the study assumed each aspirin contained 325mg).
Comparison	No aspirin.
Findings	The use of one through six aspirin per week appears to be associated with a reduced risk of a first
	myocardial infarction among women. Those 50 years of older as well as those with other coronary
	risk factors, appeared to have the greatest risk reductions.
Recommendations	Approximately 98% of the cohort was white. The self-reporting nature of the study may have
and/or Limitations	provided untrue data (over-stating or under-stating how much aspirin is taken each week). The
	dosage of 1pill of aspirin (325mg) was not provided to the participants, they were simply asked how
	many aspirin they take per week and responded accordingly.

Table 3i: Evaluation of Source and Level of Evidence

	Article 9
Reference	Raju, N., Sobieraj-Teague, M., Hirsh, J., O'Donnell M., Eikelboom, J. (2011). Effect of aspirin on
	mortality in the primary prevention of cardiovascular disease. The American Journal of Medicine,
	124(7), 621-629.
Level of Evidence	18
Research Design	Meta-analysis of nine randomized controlled trials.
Problem/Purpose	To obtain best estimates of the effect of aspirin on mortality in primary prevention.
Setting	Varied.
Sample/Population	(N=100 076 total). Patients without a history of symptomatic cardiovascular disease.
Aspirin	Varied dosages (75-500mg/day or alternate days).
Comparison	Placebo or no aspirin treatment.
Findings	Aspirin prevents deaths, myocardial infarction, and ischaemic stroke, and increases hemorrhagic
	stroke and major bleeding when used in the primary prevention of cardiovascular disease.
Recommendations	Included studies were conducted over a 30-year period.
and/or Limitations	
	Aspirin dose, duration of treatment, and lengths of follow-up were not uniform.
	Meta-analysis remains a retrospective research subject to the methodological deficiencies of the included studies.

Table 3i: Evaluation of Source and Level of Evidence

	Article 10
Reference	Raju, N., Sobieraj-Teague, M., Hirsh, J., O'Donnell M., & Eikelboom, J. (2011). Effect of aspirin on mortality in the primary prevention of cardiovascular disease. <i>The American Journal of Medicine</i> , 124(7), 621-629. doi: 10.1016/j.amjmed.2011.01.018
Level of Evidence	1b
Research Design	Randomized, double-blind, placebo-controlled trial.
Problem/Purpose	To evaluate the lowest dose of aspirin that would having a cardio-protective effect, while minimizing gastrointestinal side effects through the use of a low dose and alternate-day administration.
Setting	United States.
Sample/Population	Initially healthy women (N=39,876) aged 45 or older. Monitored for 10 years for a first major cardiovascular event (nonfatal myocardial infarction, nonfatal stroke, or death from cardiovascular causes).
Aspirin	100mg aspirin on alternate days.
Comparison	Placebo.
Findings	Aspirin did not affect the risk of myocardial infarction or death from cardiovascular causes. However, benefits were observed among the subgroup of women who were 65 years of age or older. The frequency of side effects related to bleeding and ulcers was increased among women who received aspirin.
Recommendations and/or Limitations	Reasons for any sex-based differences in the efficacy of aspirin for prevention are unclear and require further exploration. The age-based difference also requires further investigation.

Table 3k: Evaluation of Source and Level of Evidence

	Article 11	
Reference	Wolff, T., Miller, T., &Ko, S. (2009). Aspirin for the primary prevention of cardiovascular events: An	
	update of the evidence for the U.S. preventive services task force. <i>Annals of Internal Medicine</i> , 150(6),	
	405-410. doi:10.7326/0003-4819-150-6-200903170-00009	
Level of Evidence	28	
Research Design	Randomized controlled trials (RCTs), case-control, meta-analysis, and systematic reviews of aspirin	
	versus control for the primary prevention of cardiovascular disease.	
Problem/Purpose	To determine the benefits and harms of taking aspirin for the primary prevention of myocardial	
	infarctions, strokes, and death.	
Setting	Varied (including the United States, United Kingdom, and Italy).	
Sample/Population	Individuals without history of cardiovascular disease.	
Aspirin	Varying dosage from 75 mg/d to 500 mg/d. The Women's Health Study used 100 mg every other day.	
Comparison	Placebo or no aspirin.	
Findings	Aspirin reduces the risk of myocardial infarctions in men. The use of aspirin for primary prevention	
	increases the risk for major bleeding events, primarily gastrointestinal bleeds, in both men and	
	women. Men have an increased risk for hemorrhagic strokes with aspirin, although the risk is not	
	statistically significantly increased in women.	
Recommendations	The dose of aspirin used in the RCTs varied, preventing the estimation of the best dose for primary	
and/or Limitations	prevention. Several of the RCTs were performed in health professionals, potentially limiting	
	generalizability.	
	Aspirin dose, duration of treatment, and lengths of follow-up were not uniform.	

Discussion of Results

More research needs to be conducted, but preliminary evidence suggests aspirin has no benefit for primary prevention of myocardial infarction in women between the ages of 50 and 65 (Ridker, 2005). The study consisted of 39 876 initially healthy women age 45 or older who were monitored for 10 years and given 100mg of aspirin on alternate days or a placebo. The low dose used in the study by Ridker (2005) could be suggested as a potential reason why no effect was seen in regards to the reduction of MI. However, the meta-analysis conducted by Berger et al. (2006) found similarly that although aspirin reduced MI in men, it did not have a significant effect for women in reducing the incidence of MI. The same study by Ridker (2005) found that, in the subpopulation of healthy females over the age of 65, aspirin did reduce the incidence of MI. Although more research is needed, the preliminary evidence suggests that aspirin as primary prevention for MI is not recommended for those healthy women with no precipitating factors (Ridker, 2005).

There have been many studies done primarily on men in regards to aspirin and its usefulness for the primary prevention of MI. The Antithrombotic Trialists' Collaboration (2002), Bartolucci et al. (2011), Berger et al. (2006), Hayden (2002), Raju et al. (2011), and Wolff, Miller, and Ko (2009) concluded that low-dose aspirin does, in fact, decrease incidence of myocardial infarction when used as primary prevention for men. The trial done by Juul-Moller et al. (1992) concluded that low-dose aspirin in 2 035 patients aged 30 to 80 with chronic stable angina significantly reduced myocardial infarction and has been examined in some of the meta-analysis. Other trials included in the many analyses were conducted from 1988-2014; the broad range of dates makes it difficult to perform a homogenous analysis of the evidence due to the various research techniques, populations, dosages of aspirin, and other circumstances of life such as control of adjuvant medications and lifestyle factors.

Although the studies discussed above determine that aspirin does have a benefit, Ikeda et al. (2014) and Howard (2014) conclude that the current evidence is not definitive proof of aspirin's efficacy in prevention for those at low risk. In fact, the study conducted by Ikeda et al. (2014) found that low dose aspirin did not have any significant impact on reducing the risk of myocardial infarction. It should be noted, however, that this study was done on a Japanese population; this population has been shown to have a lower overall incidence of myocardial infarction risk factors such as smoking and increased body mass index (BMI), leading to a lower incidence of myocardial infarction than in white males of the Western population (Ikeda et al., 2014). The study also terminated early, was not blinded, and did not

have a placebo control due to Japanese ethical standards (Ikeda et al., 2014). These factors may have contributed to the fact that the no-aspirin group experienced a 10% increase in the uptake of aspirin by the 5th year (Ikeda et al., 2014). It is important to note that this does not mean aspirin is not effective for primary prevention, but rather more information is needed to establish substantial evidence specifically for those at low risk (Howard, 2014).

An important factor when looking at any benefit of aspirin in the primary prevention of myocardial infarction for both males and females between the ages of 50 and 65 is to examine any adverse effects of the therapy in question. In every trial and meta-analysis evaluated, the incidence of bleeding, specifically gastrointestinal bleeds and hemorrhagic stroke, increased with the use of aspirin in comparison to the control group. Although the outcome examined for this paper was solely the incidence of myocardial infarction, it is important to note that all risks and benefits must be weighed in collaboration with a health professional when deciding whether or not to take aspirin (Gaziano & Greenland, 2014).

Summary and Recommendations for Practice

Although current evidence has not provided definitive proof of the effectiveness of aspirin for the primary prevention of myocardial infarction for every individual with no previous cardiovascular disease, the preliminary evidence of this paper suggests aspirin for the primary prevention of myocardial infarction is not suitable for women aged 50 to 65, while it does hold benefits for males of the same age range (Howard, 2014). However, the evidence is not unanimous, and more research is needed before the decision can be made to recommend aspirin for primary prevention in all low-risk individuals. The U.S. Preventive Services Task Force (2009) recommendations regarding aspirin for primary prevention based on 10-year cardiovascular risk exemplifies how prescription by a physician needs to be done with each patient in conjunction with the best evidence and with consideration of the individual overall risk assessment (Gaziano & Greenland, 2014).

Nurses in acute care settings need to be educated on aspirin and the current evidence on its use for primary prevention, and made aware of the prevalence of self-medication in patient populations. The nurse has a significant role to play in education, helping each patient to assess individual risks and benefits, and in advising patients to consult their physician before self-medicating (Howard, 2014). Nurses must do a thorough medication screen with every patient, as aspirin can have many implications for the acute care setting regardless of dosage (Howard, 2014). The

increased risk of bleeding caused by aspirin can be detrimental if not taken into account before surgery and other procedures (Gaziano & Greenland, 2014; Lippincott Williams & Wilkins, 2005). For short- and long-term recommendations for nursing practice as well as their supports and barriers, refer to Table 4. Although the current evidence is not clear enough to determine definitively who

should or should not be on aspirin, the nurse is a trusted professional who does have the vital role of educating each patient about not only aspirin and its risks and benefits, but also every medication, procedure, and diagnosis.

nce is not clear enough to determine definitively who			
During the acute phase (short-term): Identification of aspirin use upon the preliminary assessment. This may be supported by the utilization of a question specifically for aspirin use on the initial assessment form along with the other current medications.			
Supports	Barriers		
Aspirin affects bleeding, which has a significant impact on many procedures in the acute period (Gaziano & Greenland, 2014). Aspirin has the potential to interact with other drugs, specifically with other anticoagulants. Identification upon first assessment helps ensure safety with medication prescription and administration. Prompt identification can ensure that screening is not forgotten in the later period of care. Chronic, low-dose aspirin administration can be a cause of gastrointestinal bleeds or hemorrhagic stroke (Gaziano & Greenland, 2014). Early identification of the cause of bleeding leads to better patient outcomes.	 The time constraints of the acute period of care can deter from a thorough preliminary assessment, specifically one including an additional question about aspirin use. The primary individual may not be conscious, or able to answer questions, or language may be a barrier. There may not be anyone with the patient who can state whether the patient is taking aspirin or not. 		
2. After the acute phase or when an opportunity presents itself (long-term): Provide education on aspirin's risks and benefits, including an individual cardiovascular risk assessment for both those individuals on aspirin for primary prevention as well as those who may benefit from aspirin for primary prevention of myocardial infarction according to current evidence (Howard, 2014). Education may be facilitated through the use of a pamphlet or brochure about aspirin. Referral to the primary physician is warranted for all individuals who are taking aspirin or who are considering taking aspirin (Howard, 2014).			
• • • • • • • • • • • • • • • • • • • •	Barriers		
decisions (Howard, 2014). Those individuals on aspirin for primary prevention may not have received the proper education about the medication and its risks and benefits, and their own risks for myocardial infarction (Howard, 2014). Those individuals who are not taking aspirin prophylactically may already be considering taking it. Proper education can lead those individuals to seeking their physician's counsel. Those individuals who are not taking aspirin for primary prevention may be at such a risk for myocardial infarction that aspirin is warranted. Proper education and screening can lead these individuals to consult their physician on the matter. Consulting with a physician leads to more informed and safe health care decisions (Howard, 2014). It also ensures that the physician is aware of any aspirin being taken by the individual and can use that information when prescribing other medications, or procedures. A pamphlet or brochure enables a person to take the information home and review it, rather than depending solely on memory of the teaching conversation with a	 The time constraints of the acute period of care can make patient education less of a priority. Lack of nurse education around current evidence surrounding aspirin for primary prophylaxis of myocardial infarction could lead nurses to not teaching their patients, or providing them with outdated information. Not all individuals a nurse sees have ready access to a physician. Not all individuals can read a pamphlet or brochure. The healthcare practitioner who takes the medication history must ask not only about prescription medications, but also over-the-counter (OTC) drugs, vitamins, supplements, traditional medicines, and herbal preparations. 		
	During the acute phase (short-term): Identification of aspin supported by the utilization of a question specifically for asp current medications. Supports Aspirin affects bleeding, which has a significant impact on many procedures in the acute period (Gaziano & Greenland, 2014). Aspirin has the potential to interact with other drugs, specifically with other anticoagulants. Identification upon first assessment helps ensure safety with medication prescription and administration. Prompt identification can ensure that screening is not forgotten in the later period of care. Chronic, low-dose aspirin administration can be a cause of gastrointestinal bleeds or hemorrhagic stroke (Gaziano & Greenland, 2014). Early identification of the cause of bleeding leads to better patient outcomes. After the acute phase or when an opportunity presents its benefits, including an individual cardiovascular risk assessme prevention as well as those who may benefit from aspirin for current evidence (Howard, 2014). Education may be facilitat Referral to the primary physician is warranted for all individualspirin (Howard, 2014). Supports Education enables patients to make informed health care decisions (Howard, 2014). Those individuals on aspirin for primary prevention may not have received the proper education about the medication and its risks and benefits, and their own risks for myocardial infarction (Howard, 2014). Those individuals who are not taking aspirin prophylactically may already be considering taking it. Proper education can lead those individuals to seeking their physician's counsel. Those individuals who are not taking aspirin for primary prevention may be at such a risk for myocardial infarction that aspirin is warranted. Proper education and screening can lead these individuals to consult their physician on the matter. Consulting with a physician leads to more informed and safe health care decisions (Howard, 2014). It also ensures that the physician is awarranted on any seprin being taken by the individual		

References

- Antithrombotic Trialists' Collaboration. (2002).

 Collaborative meta-analysis of randomized trials of antiplatelet therapy for prevention of death, myocardial infarction, and stroke in high-risk patients. *British Medical Journal*, 324(7329), 71-86. doi: 10.1136/bmj.324.7329.71
- American Society of Health-System Pharmacists (1997).

 ASHP therapeutic position statement on the use of aspirin for prophylaxis of myocardial infarction.

 American Journal of Health-System Pharmacy,
 54(17), 1984-1987.
- Bartolucci, A., Tendera, M., & Howard, G. (2011). Metaanalysis of multiple primary prevention trials of cardiovascular events using aspirin. *The American Journal of Cardiology*, 107(12), 1796-1801. doi:10.1016/j.amjcard.2011.02.325
- Berger, J., Roncaglioni, M., Avanzini, F., Pangrazzi, I., Tognoni, G. & Brown, D. (2006). Aspirin for the primary prevention of cardiovascular events in women and men: A sex-specific meta-analysis of randomized controlled trials. *The Journal of the American Medical Association*, 295(3), 306-13. doi:10.1001/jama.295.3.306
- Gaziano, J. M., & Greenland, P. (2014). When should aspirin be used for prevention of cardiovascular events?

 Journal of the American Medical Association,
 312(23), 2503-2504. doi:10.1001/jama.2014.16047
- Hayden, M. (2002). Aspirin for the primary prevention of cardiovascular events: A summary of the evidence for the U.S. preventive services task force. *Annals of Internal Medicine*, 136(2), 161-72. doi:10.7326/0003-4819-136-2-200201150-00016
- Hennekens, C., Manson, J., & Reilly, M. (2002). Who should be taking aspirin? *Patient Care*, *36*(13), 58.
- Howard, P. A. (2014). Aspirin for primary cardiovascular prevention: When is it worth the risks? *Hospital Pharmacy*, 49(6), 502-507. doi:10.1310/hpj4906-502
- Howick, J. (2009). Oxford Centre for Evidence-based

 Medicine Levels of Evidence. Accessed from

 http://www.cebm.net/oxford-centre-evidence-based-medicine-levels-evidence-march-2009/

- Ikeda, Y., Shimada, K., Teramoto, T., Uchiyama, S., Yamazaki, T., Oikawa, S., & ... Ishizuka, N. (2014). Low-dose aspirin for primary prevention of cardiovascular events in Japanese patients 60 years or older with atherosclerotic risk factors: A randomized clinical trial. *JAMA: Journal of the American Medical Association*, 312(23), 2510-2520. doi:10.1001/jama.2014.15690.
- Juul-Möller, S., Edvardsson, N., Jahnmatz, B., Rosén, A., Sørensen, S., & Omblus, R. (1992). Double-blind trial of aspirin in primary prevention of myocardial infarction in patients with stable chronic angina pectoris. The Swedish Angina Pectoris Aspirin Trial (SAPAT) Group. *Lancet*, 340(8833), 1421-1425. doi:10.1016/0140-6736(92)92619-Q
- Kappagoda, C.T., & Amsterdam, E.A. (2011). Trials of primary prevention of cardiovascular events using aspirin. *The American Journal of Cardiology*, 108(8), 1198-1200. doi: 10.1016/j.amjcard.2011.07.035
- Lippincott Williams & Wilkins. (2005). *Nursing 2005 Drug Handbook*. Ambler, PA: Author. Ed. 25, p. 355-358.
- Manson, J., Stampfer, M., Colditz, G., Willett, W., Rosner, B., Speizer, F., & Hennekens, C. (1991). A prospective study of aspirin use and primary prevention of cardiovascular disease in women. *The Journal of the American Medical Association*, 266(4), 521-527. doi:10.1001/jama.1991.03470040085027.
- Raju, N., Sobieraj-Teague, M., Hirsh, J., O'Donnell M., & Eikelboom, J. (2011). Effect of aspirin on mortality in the primary prevention of cardiovascular disease. *The American Journal of Medicine*, 124(7), 621-629. doi: 10.1016/j.amjmed.2011.01.018
- Ridker, P. M. (2005). A randomized trial of low-dose aspirin in the primary prevention of cardiovascular disease in women. *The New England journal of medicine* 352(13), 1293-1304. doi: 10.1056/NEJM0a050613
- U.S. Preventive Services Task Force. (2009). Aspirin for the prevention of cardiovascular disease: U.S. preventive services task force recommendation statement. *Annals of Internal Medicine*, 150(6), 396-404. doi:10.7326/0003-4819-150-6-200903170-00008.

- Wolff, T., Miller, T., & Ko, S. (2009). Aspirin for the primary prevention of cardiovascular events: An update of the evidence for the U.S. preventive services task force. *Annals of Internal Medicine*, 150(6), 405-410. doi:10.7326/0003-4819-150-6-200903170-00009
- World Health Organization (2015). Cardiovascular diseases (CVDs). Retrieved from http://www.who.int/mediacentre/factsheets/fs317/en/

Aspirin for Primary Prevention of Myocardial Infarction (Nemetchek)
University of Saskatchewan Undergraduate Research Journal