

**Multiple Small Feedings  
Of The Mind In  
Internal Medicine**

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# Multiple Small Feedings Of The Mind In Internal Medicine

## Objectives

- Address commonly encountered medical problems in Primary Care Practice that YOU wanted to hear about
- Provide focused information that is evidence based, practical and immediately usable
- Keep it brief and to the point – to prevent audience from falling a sleep

# **Multiple Small Feedings Of The Mind In Internal Medicine**

**1 – Women's Health**

**2 – Geriatric Medicine**

**3 - Anticoagulation**

**Multiple Small Feedings**  
**Of The Mind In**  
**Internal Medicine**

**1 – Women's Health**





"Can you give me something to make me hot"



"What a Devil! After the last flash...

# MSFM – Women's Health

How Tempted are you  
To give  
Hormone Therapy  
To that post-menopausal female patient  
Who complains every visit  
About her  
Vaso-Motor Symptoms?

# MSFM – Women's Health

## Case # 1

- 58 yo wf comes to office with vaso-motor symptoms that are affecting the quality of her life
- Hx = Stage I HTN, mild Hyperlipidemia, non-smoker
- Vitals and PE unremarkable
- She inquires about Hormone Therapy

# MSFM – Women's Health

## Case # 1

### USPSTF Guidelines for HT in Postmenopausal Women

JAMA; 2017 Dec 12; 318(22):2224-2233

US Preventative Services Task Force

# MSFM – Women's Health

## Case # 1

- Estrogen Tx given to 40% of women (1988-1994)
- Then - Women's Health Initiative (WHI) showed harm with Tx
- So - Estrogen Tx went down to 10% (2010)
- (Estrogen + Progesterone) = Reduced the risk of Fx & DM
- Then - 18 Clinical trials reviewed (since 2012 USPSTF recommendation)

# MSFM – Women's Health

## Case # 1

- HT increased the risk of – Invasive Breast Cancer, Venous Thromboembolism, CAD
- Risk >> Benefit
- This is for Post Menopausal Asymptomatics
- Not for Premature Ovarian Failure

# MSFM – Women's Health

USPSTF Recommends **AGAINST**

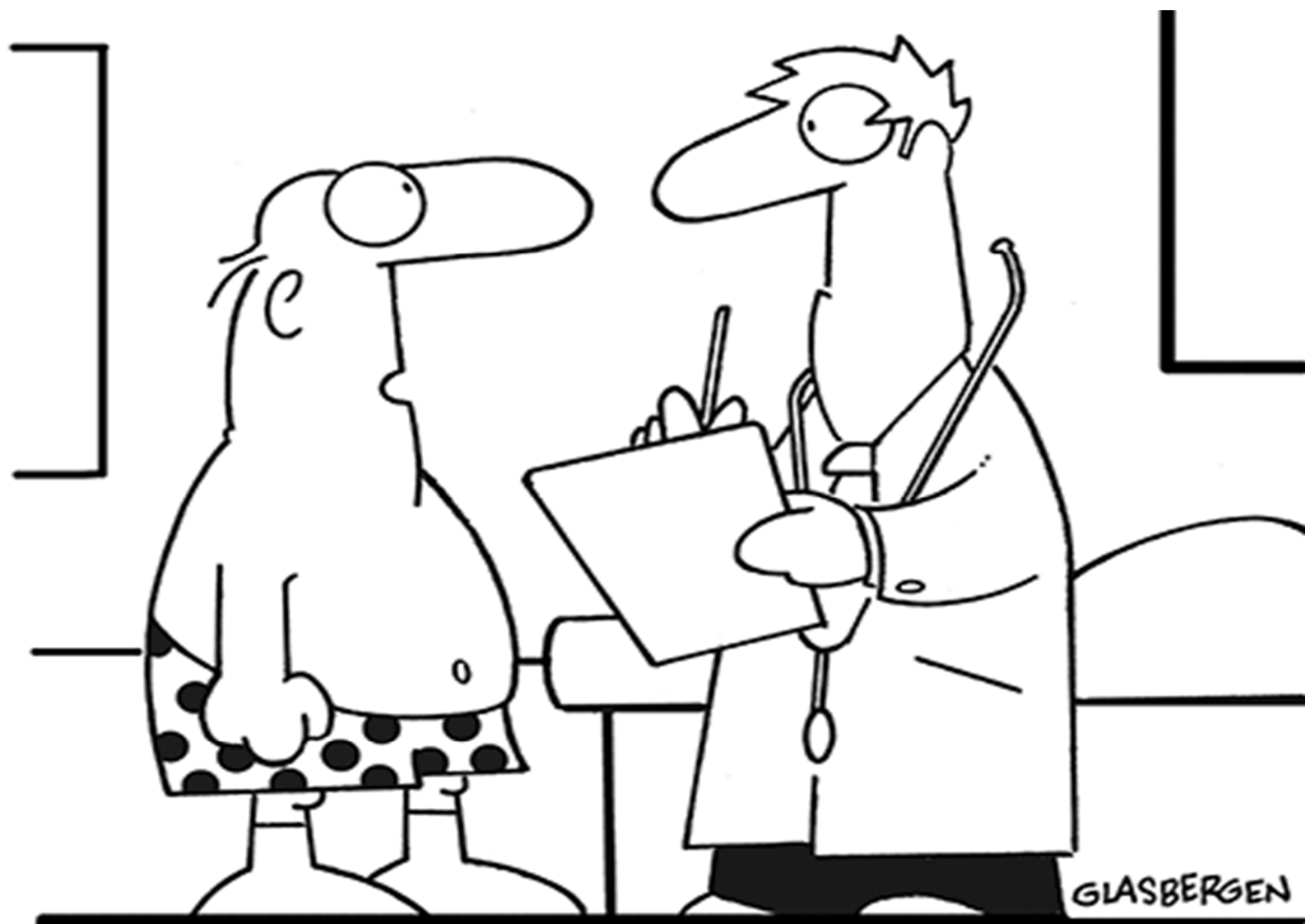
**1** - Combined Estrogen & Progesterone for the Primary Prevention of Chronic Conditions in Post Menopausal Women

**OR**

**2** - Estrogen alone for the Primary Prevention of Chronic Conditions in post menopausal women who have had hysterectomy

(Type D Recommendation for both)





## MSFM – Women's Health

How well-versed are you  
in Recommending  
Nutritional Supplements  
to your Patients?

# MSFM – Women's Health

## Case # 2

- 75 yo bf new to your practice has Hx of DJD, Osteopenia & HTN
- She is well functional and independent
- Her vitals and PE are normal
- She wants your opinion on Calcium and Vitamin D Supplementation for fracture prevention

# MSFM – Women's Health

## Association Between Calcium or Vitamin D Supplementation and Fracture Incidence in Community-Dwelling Older Adults (Systematic Review and Meta-Analysis)

JAMA. 2017 Dec 26; 318(24):2466-2482

Zhao J G, et al

# MSFM – Women's Health

## Case # 2

- 33 Randomized Trials (n = 51,145)
- Calcium OR Vit D = NO significant association with Hip Fx (Ca:- RR=1.53, Vit D:- RR=1.21)
- Calcium + Vit D = NO significant association with Hip FX (RR=1.09)
- Calcium OR Vit D (OR Combined Tx) = NO significant association with Vertebral, Non-Vertebral or Total Fx

# MSFM – Women's Health

## Case # 2 (3 older studies - contrasted)

- 1 - “Vitamin D and calcium supplementation prevents osteoporotic Fx in elderly community dwelling residents: A pragmatic population-based 3-year intervention study” (Larsen, E R et al; J Bone Mineral Res. 2004; 19:370-378)
- 2 - “A higher dose of Vitamin D reduces the risk of falls in nursing home residents: A randomized, multiple-dose study” (Broe, K E et al; J of Amer Geriatr Soc. 2007;55:234-239)

# MSFM – Women's Health

## Case # 2 (3 older studies - contrasted)

- 3 – Fracture prevention with Vitamin D supplementation. A meta-analysis of randomized controlled trials. (Bischoff-Ferrari H A, et al; JAMA. 2005; 293(18)257-2264)

# MSFM – Women's Health

## Case # 2 (3 older studies)

- Fractures decreased with Calcium & Vitamin D
- Falls Decreased with Vitamin D in NH residents  
(due to increased muscle strength in vitamin D deficient patients?)(supplement alone did not reduce falls)
- 700 iu – 800 iu of Vitamin D (NOT 400 iu) =  
Reduced hip & non-vertebral Fx in ambulatory  
OR institutionalized older adults



# MSFM – Women's Health

## Case # 2

### So ..... Use Common Sense

- Vitamin D has many “non-bone benefits”
- Vitamin D levels are lower in population due to avoidance of SUN due to risk of skin Ca
- Colon Cancer recurrence was higher after cancer Tx in patients with low Vitamin D levels
- Checking 25 OH-Vitamin D level is reasonable (keep level in mid-range = 45 - 65)
- Recommend diet, rich in calcium
- Recommend “resistance” exercises
- Remember --> Bone health begins in-utero !!!

# MSFM – Women's Health

## Case # 3

- 80 yo wf with hx of osteoporosis stable on Bisphosphonate Tx for 10 years wants to “get off” the Tx – because she heard of “side effects”
- Hx = HTN, DJD, Osteoporotic Fx 10 years ago, DM Type 2
- Asymptomatic & functional

# MSFM – Women's Health

## Case # 3

Long-term oral bisphosphonate therapy  
and fractures in older women: The  
Women's Health Initiative Study

J Amer Geriatric Soc.

Drieling R L et al; 2017: 65(9)1924-1931

# MSFM – Women's Health

## Case # 3

- Women's Health Initiative reported data
- n = 5120 (median age = 80) (f/u = 4 yrs)
- Duration of Tx (Tx group) = 3, 5, 6-9, 10-13 yrs
- Duration of Tx (control group) = 2 yrs
- Fx Risk = (10-13 yr Tx group) >> (2 yr group)
- HR = 1.30 (NO prior Fx group = Prior Fx group = Cancer group)
- Hip Fx = Vertebral Fx = Wrist Fx

# MSFM – Women's Health

## Case # 3

- ACP guideline recommends Tx of osteoporosis (with T score < 2.5) with pharmacologic Tx for **5 years** (Treatment of low bone density or osteoporosis to prevent fractures in men and women: A clinical practice guideline update from the American College of Physicians. Ann Intern Med. 2017; 166(11):818-839)
- Risk of Osteo-Necrosis of the Jaw & atypical femoral Fx increase with duration of Tx
- Limiting Bisphosphonate Tx to 5 yrs or less is recommended



# MSFM – Women's Health

Cleanliness  
Is Next to Godliness

BUT

It has its Price

# MSFM – Women's Health

## Case # 4

- 60 yo bf comes with chronic progressive SOB
- Hx = HTN, DJD, non-smoker, no asthma
- Occupation = House cleaning
- Vitals, EKG, CXR, Pulse Ox (RA) = normal
- Exercise Stress Test = SOB, no ST-T changes
- PFT = FEV1 75 % of predicted (no reversibility)



# MSFM – Women's Health

## Case # 4

Cleaning at home and at work in  
relation to lung function decline and  
airway obstruction

Amer J Respir Crit Care Med.

ePub; 2018 Feb 16; Svanes, et al

# MSFM – Women's Health

## Case # 4

- $n = 6230$
- Women cleaning at home & at work
- 20 year Cohort Study
- 22 Study centers
- At least 1 Lung Function Study done

# MSFM – Women's Health

## Case # 4

- Women exposed to “cleaning chemicals” at work OR at home had more rapidly declining FEV1, FVC (-22.1,  $p=0.01$ )
- Cleaning sprays & liquids had similar effects
- Men were NOT affected
- No significant association with COPD
- i.e. -- ASK ABOUT OCUPATIONAL EXPOSURE TO CLEANING CHEMICALS OR ENVIRONMENTAL POLLUTIONS IN WOMEN WITH DIMINISHED LUNG FUNCTION AND NO HX OF SMOKING OR ASTHMA

**Multiple Small Feedings**  
**Of The Mind In**  
**Internal Medicine**

**2 - Geriatrics**

RETIREMENT  
HOME

SPEED  
DATING



# MSFM – Geriatrics

When you see a patient in the office or the hospital, you stratify their health risk from

**“Medical Perspective”**

HAVE YOU EVER

Thought of looking at them from the  
Point of view of

**“Financial Risk?”**

(Should This Be The 5<sup>th</sup> Vital Sign Instead of The “Pain”?)



**“When you review your retirement fund then**

# MSFM – Geriatrics

## Case # 1

### Wealth-Associated Disparities in Death and Disability in the United States and England

JAMA Intern Med; 2017 October 23

Makaroun L K, et al



# MSFM – Geriatrics

## Case # 1

- Examined Relationship between WEALTH & MORTALITY (not just “current income”)
- Compared age < 65 & >65 (i.e. access to government sponsored health insurance, OR access to care)

# MSFM – Geriatrics

## Case # 1

- $n = 12,173$  (USA),  $n = 7,599$  (UK)
- Age Stratification = (54 – 64) vs. (66 – 76)
- As WEALTH decreased → DEATH & DISABILITY increased (over 10 yrs & across the quintiles)
- Wealth of < \$39,000 = 17% Mortality Risk (HR = 3.3), 48% Disability Risk (HR = 4.0)
- Wealth of > \$560,000 = 5% Mortality Risk, 15% Disability Risk

# MSFM – Geriatrics

How Successful are you in  
Getting your Elderly  
Patients

OFF Of Their  
Sleeping Pills?

(Are you kidding me? – That's a joke ... right?)



# MSFM – Geriatrics

## Case # 2

- 88 yo wf comes with her daughter who wants to discuss functional decline and future care planning
- Hx – mild Dementia, HTN, DJD, COPD
- Medications – Zolpidem 10 mg HS x 15 Yrs. (+ others)

# MSFM – Geriatrics

## Case # 2

The Use of Benzodiazepines Receptor  
Agonists (BZRA) and the risk of  
Hospitalization for Pneumonia: A  
nationwide Population-based nested case-  
controlled Study

CHEST. 2017;153:161-171

Chen T, et al

# MSFM – Geriatrics

## Case # 2

- n = 12,002 (between 2002 – 2012)
- Risk of Hospitalization for Pneumonia =  
BZRA use (OR = 1.86), Benzo use for Sleep (OR = 2.42), Benzo use for Anxiety (OR = 1.53), Non-Benzo Hypnotic Agent use (OR = 1.60), Midazolam use (OR = 5.77)
- Increased risk of MVA with use (FDA Announcement: 2013; Jan 10)

## MSFM – Geriatrics

“What can I get you in your Coffee Sir?”

“I’ll have Cream and Sugar,

BUT

My Dad will take

a pinch of Methylphenidate,

Please”



10-THIRDS OF SENIORS HAVE TROUBLE SLEEPING, SAYS ST

FRED HAS TROUBLE  
SLEEPING AT NIGHT, AND  
AND HIS DOCTOR CAN'T  
FIGURE OUT WHY!

STAYS  
TROUBLE  
SERVICE



# MSFM – Geriatrics

## Case # 3

### Methylphenidate for Apathy in Community-Dwelling Older Veterans with mild Alzheimer's Disease: A Double-Blind Randomized Placebo-Controlled Trial

Am J Psychiatry. 2017; Sept 15;

Padala R R et al

# MSFM – Geriatrics

## Case # 3

- n = 60 !!!, (Duration = 12 weeks)
- Mean Age = 77 yrs
- Evaluation at 4, 8, 12 weeks
- Measured by “Clinical Global Impression Scale (CGI)
- Improved = Apathy, Depression, Functional Status, Cognition ... & thus ..... care-giver burden

# MSFM – Geriatrics

So.....

Now with Opioid Crisis

Are you Switching  
Your “Arthritic Patients”  
To NSAIDs?

WELL, AT LEAST  
WE WERE  
**HALF** RIGHT!..

~~**PAIN**~~

FDA

**vioxx:**  
**KILLER**

FDA APPROVED

MERCK  
CO. INC.

**vioxx**  
USERS





# MSFM – Geriatrics

## Case # 4

The Risk of Major NSAID Toxicity with  
Celecoxib, Ibuprofen or Naproxen: A  
Secondary Analysis of the PRECISION  
Randomized Controlled Clinical Trial

Am J Med. 2017; July 26;

Solomon, D H et al;

# MSFM – Geriatrics

## Case # 4

- n = 24,081 (with OA & RA)
- All with Moderate CV Risk & Symptomatic Arthritis
- Tx with Celecoxib **OR** Ibuprofen **OR** Naproxen
- All received PPI
- Secondary Analysis of PRECISION Trial



# MSFM – Geriatrics

## Case # 4

- 5% = Major Toxicity over 1-2 yrs
- Toxicity = CV events, GI bleed, Kidney Injury & All Cause Mortality
- Toxicity = Celecoxib (4.1%), Ibuprofen (5.3%), Naproxen (4.8%)
- After adjusting for ASA and Geographic Region  
→ Ibuprofen (41% higher) & Naproxen (19% higher) Risk → THEN → Celecoxib Risk

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**3 - Anticoagulation**

YOU CAN'T GET THE  
GOOD STUFF ANYMORE...  
I SAW THIS DAY  
COMING WHEN THEY  
INVENTED BLOOD  
THINNERS.



# MSFM – Anticoagulation

Just When you Thought That

ASA was for Arterial

And

Anticoagulation was for Venous

“Clogged Plumbing” .....

Than comes This .....

# MSFM – Anticoagulation

## Case # 1

### Aspirin or Rivaroxaban for VTE Prophylaxis after Hip or Knee Arthroplasty

N Engl J Med. 2018 Feb 22;378:699-707

Anderson D R et al;

# MSFM – Anticoagulation

## Case # 1

- n = 3424 (almost Equal Total Hip & Total Knee Sx)
- All Received Rivaroxaban 10 mg Post-Op x 5 d
- Then Randomized
- Aspirin 81 mg OR Rivaroxaban 10 mg
- Total Tx = 14 d for Knee & 30 d for Hip
- NO difference in VTE at 90 d (0.06% vs 0.07%)
- NO difference in Major Bleed (0.04% vs 0.03%)
- No difference in Clinically Important Bleed (1.2% vs 1.0%)

# MSFM – Anticoagulation

## What About Cancer Associated VTE?

(where Low-Molecular Wt Heparin in past was  
Superior to Vitamin K Antagonists, Creating  
Logistical problem in Cancer Patients)

# MSFM – Anticoagulation

## Case # 2

# Edoxaban for the Treatment of Cancer-Associated Venous Thromboembolism

N Engl J Med. 2018 Feb 15;378:615-624

Raskob G E et al;



## MSFM – Anticoagulation

- n = 1046 ( Modified Intention-to-Tx Analysis)
- Low Molecular Wt Heparin x 5 days first
- Then Edoxaban 60 mg/d vs. Deltaparin 200 iu/Kg/d for 30 d followed by 150 iu/Kg/d
- Duration = 6 – 12 months
- Recurrent VTE (+) Major Bleed (as follows)
- Edoxaban = 12.8% vs. Deltaparin (11.3%)
- VTE Alone = Edo (7.9%) vs. Heparin (11.3%)
- Major Bleed = Edo (6.9%) vs. Heparin (4.0%)

# MSFM – Anticoagulation

If You can Give ASA  
for VTE Prevention after Hip & Knee Sx  
(=Venous Dz)

Can you give NOACs for PVD?  
(=Arterial Dz)

falls, you take a baby aspirin.



# MSFM – Anticoagulation

## Case # 3

### Rivaroxaban with or Without Aspirin in Patients with stable Peripheral or Carotid Artery Dz

Lancet; 2018 Jan 20; Anand, Bosch et al;

# MSFM – Anticoagulation

## Case # 3

- n = 7470 (558 centers)
- Rivaroxaban (5mg BID) + ASA vs. ASA alone
- Combined CV Death + MI + CVA (as follows)
- R + A = 5% vs. ASA = 7% (HR=0.72, p=0.0047)
- Major Adverse Amputation (as follows)
- R + A = 1% vs. ASA = 2% (HR=0.54, p=0.037)
- Major Bleed (as follows)
- R + A = 3% vs. ASA = 2% (HR=1.61, p=0.0089)

# MSFM – Anticoagulation

## Case # 3

- Rivaroxaban + ASA slightly better than ASA alone or Rivaroxaban alone
- BUT – Combination increased Bleeds
- Rivaroxaban 5 mg BID = ASA alone  
(Not yet for main stream ... Cost?)



**"I'm going to prescribe something that works like aspirin but costs much, much more"**

# MSFM – Anticoagulation

“Thinner the Blood Better it is”

OR

IS IT?

What about  
NOACs and Mortality from Intra-  
Cranial Hemorrhage (ICH)?



# MSFM – Anticoagulation

## Case # 4

Association of Intracranial Hemorrhage  
among patients taking Non-Vitamin K  
Antagonists vs. Vitamin K Antagonists Oral  
Anticoagulants with In-Hospital Mortality

JAMA. 2018;319(5):463-473

Inohara T et al;

# MSFM – Anticoagulation

## Case # 4

- Retrospective Cohort Study (n = 141,311)
- In-Hospital Mortality Assessed
- Warfarin vs. NOACs vs. 1 or 2 Anti-Plt drugs
- Mortality = W (32.6%), N (26.5%), AP (22.5%)
- Adjusted Risk Difference = N < W (by 5.7%)

# MSFM – Anticoagulation

When was the last time  
You saw a  
Patient with Pulmonary Embolism  
Present with the Syncope?

(Of Course there are people who consider PE as  
differential Dx for everything including “Baldness”!)



**“She’s going to need a prescription for**

# MSFM – Anticoagulation

## Case # 5

### Prevalence of Pulmonary Embolism in Patients with Syncope

JAMA Intern Med; ePub 2018 Jan 29;

Costantino, et al

# MSFM – Anticoagulation

## Case # 5

- Prospective Observational Study
- Database from US, Canada, Denmark & Italy
- n = 1,671,944 (with Syncope) (PE as follows)
- Prevalence = 0.06% - 0.55% (all patents)
- Prevalence = 0.15% - 2.10% (hospitalized Pt)
- At 90 days = 0.14% – 0.83% (all Pt)
- At 90 days = 0.35% - 2.63% (hospitalized Pt)
- Hospitalization INCREASED VTE risk by 3X

## **Multiple Small Feedings Of The Mind** **In Internal Medicine**

Remember .....

Most things you know today will be  
obsolete in the near future ...

So ....

Everything, including our thinking must  
change and evolve with time

# Multiple Small Feedings Of The Mind In Internal Medicine

Confucius once said about  
The Change:

Politicians and diapers should be  
changed regularly

And

Both for the SAME reason



**Multiple Small Feedings Of The Mind**  
**In Internal Medicine**

Unfortunately .....

We have only 2 choices  
for the Presidedency

While 50 for Miss America

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