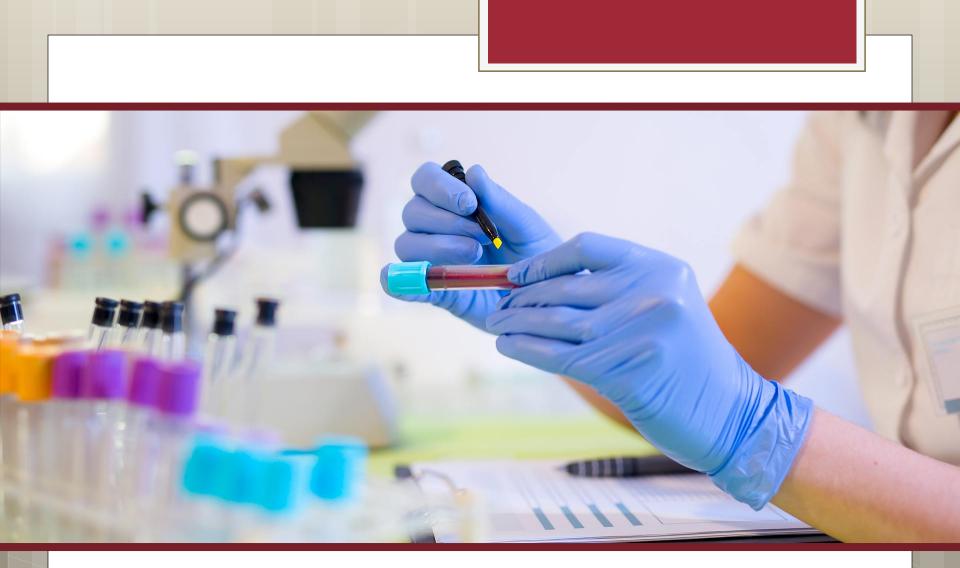
An evaluation of warfarin usage of an urban district level hospital in KwaZulu-Natal

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Introduction

Warfarin is an inexpensive and commonly used drug that is regularly prescribed in the public healthcare sector

Usage of this drug is **labour intensive** and managing patients on warfarin is complex.



Problems associated with warfarin:

- Drug interactions
- Drug-food interactions
- Life threating complications due to sub therapeutic or excessively elevated INR's
- o Costs
- Quality of life



Study Aim

To evaluate **the quality of care** of patients on warfarin therapy presenting to an urban district hospital in KwaZulu-Natal, by assessing the **financial** and **psychosocial** burden of treatment

Study Setting: **Wentworth** Hospital, eThekwini district, KwaZulu-Natal

Study Design: **Cross sectional**, observational, analytical



Three components:

1. <u>A structured patient interview</u>

 Quality of care, patient expenses and patients' willingness to pay for alternatives

2. <u>Six-month retrospective outpatient file review</u>

 Demographics, clinical profile of the patients, ascertaining outpatient costs and finding objective evidence of warfarin related hospital admissions

3. Six month retrospective inpatient file review of any admissions

• Cost determination

- The participants' outpatient and inpatient files were retrospectively reviewed for the preceding **six-month period**
- All costs involved with each warfarin related outpatient visit and inpatient stay were calculated.
- The maximum time for a follow up appointment for these patients is four weeks, hence the sampling included all patients currently on warfarin therapy from the hospital.

Inclusion criteria:

Patients attending the **Friday clinic** were included in the study as well as patients admitted for complications arising out of their warfarin usage. Patients who were **admitted** after hours were also included in the study and their clinical files were retrieved from the admission ward.

Exclusion criteria:

- 1. Patients presenting for routine INR monitoring to the medical outpatient department on any other day
- 2. Patients with lost clinical files. A patient's file was deemed to be lost if it could not be found on more than two separate occasions
- 3. Patients on treatment for less than three months

Results

A total of **135** patients were booked for the clinics for that specific time period

A total of **128** patients were eligible for the study of which 18 (14%) were excluded (nine were on treatment for less than three months and nine patient's files were not found)

Final participant number of **110** patients.

Quality of care

	Mean (SD)	Median	IQR
Treatment satisfaction	3,89 (1,03)	4	(4-5)
I am often frustrated with having to come to the hospital every month for blood tests	3,75 (1,22)	4	(2-5)
I sometimes wish that I did not have to take warfarin	4 (1,18)	4	(3-5)
Willing to pay for an alternative drug	3,82 (0,84)	4	(2-4)

Table 3 Quality of care (N=110)

1= strongly disagree, 2= disagree, 3= neutral, 4= agree, 5= strongly agree

Table 1: Relationship between age and variables measured

		Age		Total	P- values
		<60	>60		vulueb
		(n = 53)	(n = 57)		
Sex		N (%)	N (%)		
	Male	24 (45,3%)	26 (45,6%)	50	0,9
	Female	29 (54,7%)	31 (54,4%)	60	
Indication					
	AF	9 (17,0%)	33 (57,9%)	42	< 0.001
	Prosthetic heart valve	33 (62,3%(14 (24,6%)	47	< 0.001
	DVT	5 (9,4%)	4 (7,0%)	9	0,6
	PE	3 (5,7%)	1 (1,8%)	4	0,4
	Other	4 (7,5%)	5 (8,8%)	9	0,8
Co-Morbidities					
	Diabetes	6 (11,3%)	17 (29,8%)	23	0,02
	Prosthetic heart valve with co- morbid AF	9 (17,0%)	7 (12,3%)	16	0,5
	Hypertension	16 (30,2%)	40 (70,2%)	56	< 0.001
	Dyslipidaemia	4 (7,5%)	14 (24,6%)	18	0,02
	Ischaemic heart disease	1 (1,9%)	10 (17,5%)	11	0,009
	Cardiac Failure	3 (5,7%)	15 (26,3%)	18	0,003
	COPD	1(1,9%)	3 (5,3%)	4	0,6
	Asthma	3 (5,7%)	3 (5,3%)	6	0,9
	Other	24 (45,3%(19 (33,3%)	43	0,2
Number of comorbidities					
	0	17 (32,1%)	6 (10,5%)	23	0,005
	1	14 (26,4%)	12 (21,1%)	26	0,508
	2	17 (32,1%)	15 (26,3%)	32	0,506
	>=3	5 (9,4%)	24 (42,1%)	29	< 0.001
Time spent at hospital					
nospital	0 - <5 hours	16 (30,2%)	13 (22,8%)	29	0,38
	\geq 5 - <6 hours	14 (26,4%)	32 (56,1%(46	0,002
	≥ 6 hours	23 (43,4%)	12 (21,1%)	35	0,012
Willingness to pay					
	Not willing to pay	9 (17,0%)	8 (14,0%)	17	0,669
	R0-R49	16 (30,2%)	28 (49,1%)	44	0,04
	R50-R99	17 (32,1%)	15 (26,3%)	32	0,506
	R100-R199	8 (15,1%)	3 (5,3%)	11	0,086

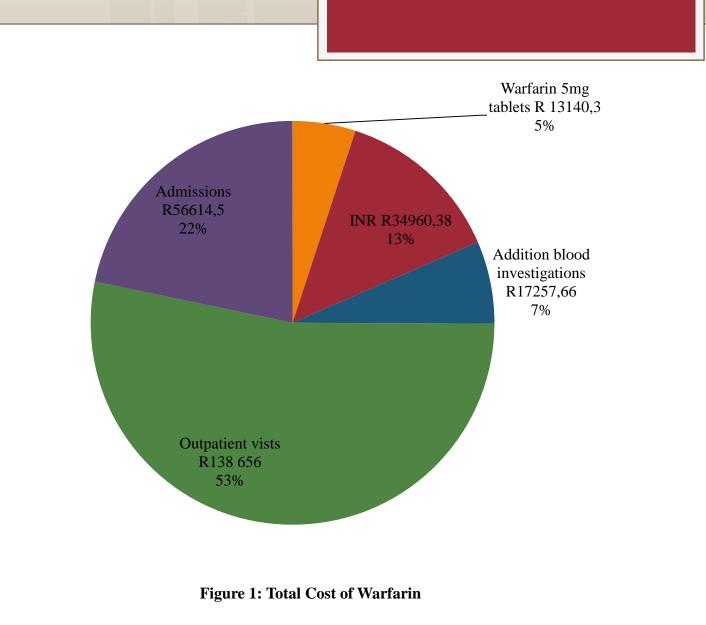
Results

84,5% willing to pay for alternative drug

AF= Atrial Fibrillation, DVT= Deep Vein Thrombosis, PE= Pulmonary Embolism, COPD= Chronic obstructive Pulmonary Disease

		Ν	ZAR	IQR/SEM
Total costs per patient per month	Median (IQR)	110	295.05	(283.42 - 333.17)
	Mean (SEM)	110	394.89	(53.03)
Non-valvular AF costs per	Median (IQR)	42	294.40	(283.85 - 345.10)
patient per month	Mean (SEM)	42	430.54	(116.14)
Other Indications per patien	Median (IQR)	68	296.13	(279.97 - 330.2)
per month	Mean (SEM)	68	372.87	(47.69)
Inpatient and	Median (IQR)	4	12 141.00	(5385.13 - 22922.13)
emergency department cost per patient	Mean (SEM)	4	14 153.63	(5866.00)
Outpatient cost per patient	Median (IQR)	110	1 764.80	(1690.83 - 1982.22)
	Mean (SEM)	110	1 854.68	(46.37)

Table 2: Costs of prescribing Warfarin



Main findings:

- 1. Warfarin impacts the **quality of life** of our patient population, to the extent that patients would be willing to pay for an alternative drug
- 2. Warfarin, although expensive, is a **cost effective** treatment option in our setting

Additional findings:

- 1. Having a specific clinic day and a **dedicated** warfarin clinic improves quality of care
- 2. Atrial fibrillation is a disease of the elderly and that prosthetic heart valves are more prevalent in those under 60 years of age.
- 3. There is a statistical significant prevalence of **hypertension** in those over 60, which can be linked to the increased incidence of atrial fibrillation in the same age group

- 4. Frequency of clinic visits and admission costs are main drivers of total costs
- DOACs, although superior, appear to be too expensive at this stage
- 6. Estimated public sector cost:
 - Dabigatran: R673.85 and R700.80 per month
 - Rivaroxaban between R545.96 and R818.94 per month

But would the DOACS be worth it?

• Pro's:

- Routine drug monitoring is not recommended
- More effective
- +/-Similar major bleeding risk
- Less drug and food interactions

o Con's

- No regularly available antidote
- Not suitable in renal failure
- Contraindicated in mechanical heart valves



Conclusion

Warfarin has an impact on our patient's **quality of life**, but it is still the most **cost effective** anti-coagulant in our setting.

DOACs will only be a cost effective alternative if the current estimated public sector drug **price** for both Rivaroxaban and Dabigatran are approximately **halved**.

Study limitations

- Missing files
- Paper filing system
- Limitations of a cross sectional study design
- Time frame
- Difficulties in standardizing costs

Further recommendations

We suggest a prospective, **head to head trial** that compares both cost effectiveness and quality of life of patients on warfarin and DOACs at a district health level.

Ethical Considerations

Permission for this study was obtained from the University of KwaZulu-Natal's Biomedical Research Ethics Committee, BE364/17, the Department of Health's ethical review board, KZ_2017RP4_664, and the CEO of Wentworth Hospital. Written informed consent was individually obtained from every patient included in this study.

Contributions and Acknowledgements

Author contributions:

Dr Laäs was the principal investigator and Prof M Naidoo was the research supervisor

Conflicts of interests: None

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