Cost and Length of Stay Associated with Vancomycin-Induced Nephrotoxicity

Vanessa Stevens, PhD¹; Minkyoung Yoo, PhD¹; Jack Brown, PharmD^{2,3}

¹Pharmacotherapy Outcomes Research Center, University of Utah; ²University of Rochester Medical Center; ³Wegmans School of Pharmacy at St. John Fisher College

<u>vanessa.stevens@pharm.utah.edu</u>

Abstract Number PIN51

Abstract

Objectives: Decreased susceptibility to vancomycin has led to recommendations for increased serum concentration targets for some infections, which has been shown to increase the risk of nephrotoxicity (NT) [1-3]. The objective of this study was to determine if vancomycin-induced nephrotoxicity has a significant impact on cost from the hospital perspective or on length of stay. **Methods:** A cost of illness study from the hospital perspective. We conducted a secondary analysis of a cohort of 398 randomly selected inpatients receiving vancomycin in a tertiary care hospital in Rochester, New York, USA. Total and variable costs were generated by hospital accountants using micro-costing methods. NT was defined as 0.5 or 50% or greater increase in serum creatinine from baseline. Generalized linear models with log link and gamma distribution and semi-log regression were used to model total and variable costs and length of stay, respectively. Cost estimates are reported in 2009 USD.

Results: 49 (12%) of patients had NT. The unadjusted median variable costs for patients with NT were higher than for patients without NT (\$47,511 vs. \$22,355, p<.0001). On multivariable analysis, variable costs were 17% greater for patients with NT compared to patients without, but this difference was not statistically significant. The median length of stay for patients with NT was two-fold greater than for patients without NT (22 vs. 11 days, unadjusted, p<.0001). After accounting for severity of illness and other factors, NT patients stayed on average 46% longer than non-nephrotoxic patients.

Conclusions: Patients with NT have significantly increased length of stay relative to patients without NT. Further research is needed to confirm whether increased length of stay has a meaningful impact on costs. As vancomycin susceptibility continues to decrease, higher doses of vancomycin may lead to an increased incidence of NT and an increase in resource utilization among these patients.

Methods

Database Design

Secondary analysis of a retrospective cohort study of adult inpatients at University of Rochester Medical Center (URMC) from January 1, 2008 – December 31, 2009. Patients were eligible if:

- 18 or older on admission
- Non-neutropenic (Baseline ANC > 1000 cells/mm3)
- Received 2 or more days of IV vancomycin
- Baseline serum SCr < 2.0 mg/dL

Patients were excluded if they had cystic fibrosis, did not have a serum V concentration drawn within 96 hours of V initiation, or had fewer than 2 SCr values drawn

Study Design and Definitions

- Cost of illness and length of stay (LOS) study from the hospital perspective.
- Total and variable costs generated by hospital accountants using micro-costing methods.
- Costs were adjusted to 2009 USD using the Personal Consumption Expenditures (PCE) Hospital and Nursing Home Services index

Analysis

- Baseline characteristics for patients with and without NT were compared
- Generalized linear models (GLM) with log link and gamma distribution were used to model total and variable costs as a function of NT
- Semi-log regression was applied to model LOS as a function of NT controlling for other covariates

Conclusions

- The median LOS for NT patients was twice that of non-NT patients
- Adjusted costs appeared to be greater in the NT group, but the difference was not significant

Limitations

- Differences in micro-costing methods across institutions may limit generalizability
- Relatively small sample size

References

- 1. Lodise TP, Lomaestro B, Graves J, Drusano GL. Larger Vancomycin Doses (at Least Four Grams per Day) Are Associated with an Increased Incidence of Nephrotoxicity.
- Antimicrob Agents Chemother 2008; 52(4): 1330-6.
 Lodise Thomas P, Patel N, Lomaestro Ben M, Rodvold Keith A, Drusano George L. Relationship between Initial Vancomycin Concentration-Time Profile and Nephrotoxicity among Hospitalized Patients. Clinical Infectious Diseases 2009; 49(4): 507-14.
- 3. Patel N, Pai MP, Rodvold KA, Lomaestro B, Drusano GL, Lodise TP. Vancomycin: We Can't Get There from Here. Clinical Infectious Diseases **2011**; 52(8): 969-74.

Results

 Table 1. Baseline Characteristics

Variable	NT	No NT	
n (%)	(n=49)	(n=349)	р
Hospital Costs; Mean (SD)			
Total Costs (TC)	74514 (73104)	48837 (82659)	<.0001
Variable Costs (VC)	45574 (44178)	30129 (50737)	<.0001
Fixed Costs (FC)	28940 (29966)	18708 (32469)	<.0001
Length of Stay; Mean (SD)	28 (20)	20 (28)	<.0001
Age; Mean (SD)	57 (57)	58 (17)	<.0001
Male	28 (57)	204 (58)	0.86
Race			
Black	8 (16)	33 (10)	0.06
White	40 (82)	276 (79)	
Other	1 (2)	40 (11)	
NT Grade (N=49)			
Mild	12 (24)	_	
Moderate	25 (51)	_	
Severe	12 (25)	_	
C. difficile Infection	1 (2)	13 (4)	0.55
Developed Need for Dialysis	12 (24)	3 (1)	<.0001
APACHE II Score; Mean (SD)	13 (6)	11 (6)	0.01
CCI; Mean (SD)	3 (2)	2 (2)	0.02
Any Major Procedure	22 (45)	153 (44)	0.89
Any Minor Procedure	45 (92)	300 (86)	0.26
Admitted to ICU	27 (55)	150 (43)	0.11
Admission due to Infection	17 (35)	162 (46)	0.12
Vancomycin Indication			
Blood Stream Infection	6 (12)	21 (6)	0.26
Empiric	34 (70)	265 (76)	
Other	9 (18)	63 (18)	
Died in Hospital	18 (37)	35 (10)	<.0001

Figure 1. Mean and Median Total Hospital Costs and LOS for patients with and without NT

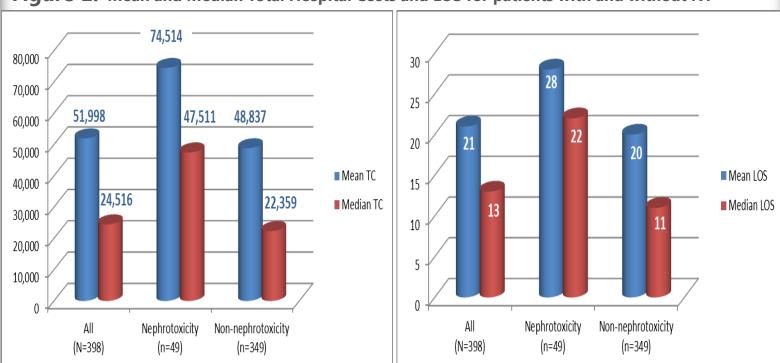


Table 2. Results from GLM cost model and Semi-log LOS regression (N=398)

Variable Coeff (SE)	Total Hospital Costs^	LOS#
Nephrotoxicity	0.17 (0.15)	0.46* (0.11)
Died in Hospital	-0.42* (0.14)	-0.67* (0.11)
Developed Need for Dialysis	0.66* (0.26)	0.20 (0.20)
APACHE II Score	0.04* (0.01)	0.03* (0.01)
CCI	0.06* (0.02)	0.04* (0.02)
DRG Weight	0.03* (0.01)	0.03* (0.0)
Any major procedure	0.61* (0.09)	0.30* (0.07)
Any minor procedure	0.96* (0.13)	0.68* (0.10)
Admitted to ICU	0.39* (0.10)	0.11 (0.08)
Admission due to infection	-0.46* (0.09)	-0.32* (0.07)
Constant	9.09* (0.30)	1.47* (0.22)

[^] Generalized Linear Model

This work was supported by a grant (PI: J Brown) from Cubist Pharmaceuticals

[#]Semi-log regression

^{* =} p < .01