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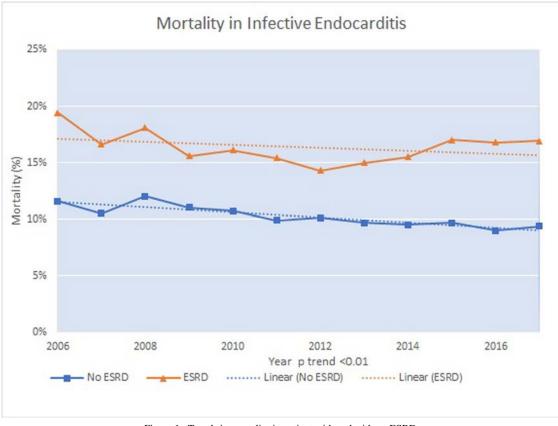


Figure 1. Trends in mortality in patient with and without ESRD.

Limitations of our study include lack of proper validation studies of ICD codes for IE. Traditionally ESRD are sicker, although we did adjust for some comorbidities, adjustment for all known and unknown confounders in a retrospective analysis is not possible. NIS has the advantage of being a large database.

In conclusion, we noticed that mortality in IE with ESRD is decreasing, albeit slower than patient with no ESRD.

Disclosures

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- Maraj S, Jacobs LE, Maraj R, Kotler MN. Bacteremia and infective endocarditis in patients on hemodialysis. *Am J Med Sci* 2004;327: 242–249.
- 2. Khan MZ, Munir MB, Khan MU, Khan SU, Benjamin MM, Balla S. Contemporary trends in native valve infective endocarditis in United States (from the National Inpatient Sample Database). *Am J Cardiol* 2020;125:1678– 1687.
- Chaudry MS, Carlson N, Gislason GH, Kamper AL, Rix M, Fowler VG, Torp-Pedersen C, Bruun NE. Risk of infective endocarditis in patients with end stage renal disease. *Clin J Am Soc Nephrol* 2017;12:1814–1822.
- 4. AHRQ. National Inpatient Sample. *Healthc Cost Util Proj* 2006. Available at: https:// www.hcup-us.ahrq.gov/nisoverview.jsp. Accessed December 2020.
- 5. Lalani T, Chu VH, Park LP, Cecchi E, Corey GR, Durante-Mangoni E, Fowler VG, Gordon D, Grossi P, Hannan M, Hoen B, Munõz P, Rizk H, Kanj SS, Selton-Suty C, Sexton DJ, Spelman D, Ravasio V, Tripodi MF, Wang A. In-hospital and 1-year mortality in patients undergoing early surgery for prosthetic valve

endocarditis. *JAMA Intern Med* 2013;173: 1495–1504.

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Burden of Thrombotic Events in Coronavirus Disease-19 (COVID-19) Patients and Effect on Outcomes (from a Multicenter Electronic Health Record Database)



Coronavirus disease-19 (Covid-19) patients have been postulated to have high risk of thrombotic events (TE). There are variable estimates of the rate of TE with COVID-19 and derived from hospital case series mostly.^{1,2} Data on unfavorable outcomes, including mortality in Covid-19 patients with TE are limited. We use a large electronic health record (EHR) database to investigate these questions.

	Be	fore propensity matching		
	COVID-19 with throm botic events (N=4140)	COVID-19 without thrombotic events (N=116375)	Risk ratio (CI)	p Value
Mortality	15.3% (633)	2% (2358)	7.55 (6.96-8.19)	< 0.001
Hospitalization	58.1% (2406)	16.0% (18632)	3.63 (3.53-3.74)	< 0.001
Mechanical Ventilation	16.9% (698)	2.1% (2416)	8.12 (7.51-8.78)	< 0.001
ARDS	18.7% (774)	1.6% (1895)	11.48 (10.62-12.41)	< 0.001
Hemodialysis	8% (330)	0.7% (850)	10.91 (9.65-12.35)	< 0.001
CRRT	2.6% (109)	0.2% (178)	17.21 (13.59-21.8)	< 0.001
	Af	ter propensity matching*		
	COVID-19 with thrombotic events (N=4140)	COVID-19 without thrombotic events(N=4140)	Risk ratio (CI)	p Value
Mortality	15.3% (633/4140)	4.8% (198/4140)	3.20 (2.74-3.73)	< 0.001
Hospitalization	58.1% (2406/4140)	25.1% (1039/4140)	2.32 (2.18-2.46)	< 0.001
Mechanical Ventilation	16.9% (698/4140)	3.9% (163/4140)	4.28 (3.63-5.05)	< 0.001
ARDS	18.7% (774/4140)	2.7% (111/4140)	6.97 (5.74-8.47)	< 0.001
Hemodialysis	3.6% (139/3861)	0.6% (26/4065)	5.63 (3.71-8.54)	< 0.001
CRRT	2.1% (85/4062)	0.3% (13/4133)	6.65 (3.72-11.91)	< 0.001

Notes: *= adjusted for age, race and gender.

Our clinical cohort included patients ≥ 18 years of age diagnosed with COVID-19 between January 20, 2020 and September 10, 2020. Covid-19 patients were identified via real-time search and analysis of more than 49 million patients from 33 healthcare organizations participating in a global health research network called TriNetX (Cambridge, MA). COVID-19 diagnosis was confirmed using specific COVID-19 diagnosis criteria recommended by the World Health Organization and Centers for Disease Control. COVID-19 patients were divided into two cohort (TE / no TE), based on presence of TE within 30 days of COVID-19 diagnosis. Validated ICD-10 diagnosis codes were used to identify TE (acute myoinfarction-AMI, pulmonary cardial embolism- PE, cerebro-vascular accident - CVA, peripheral arterial thrombosis, and deep venous thrombosis-DVT). Propensity score matching was performed for age, gender and race. Primary outcome tested was 30-day all-cause mortality post COVID-19 diagnosis. Secondary outcomes included hospitalization, need for mechanical ventilation, hemodialysis, continuous renal replacement therapy-CRRT, acute respiratory distress syndrome-ARDS and acute kidney injury. All statistical analyses were performed using TriNetX with standard methodology previously reported.³

A total of 120, 515 COVID-19 patients were identified including 4140 with TE (3.4%) – 1526: AMI, DVT – 1530, PE: 1140, CVA - 1014, peripheral arterial thrombosis: 140. Patients with TE were more likely to be male (p < 0.001), black (p < 0.001) and older (63.6 \pm 15.9 vs 47.1 \pm 18.7, p < 0.001). Table 1 shows the outcomes before and after propensity matching. COVID-19 patients with TE had higher mortality (15.3% vs 2.0%), hospitalization (58.1% vs 16.0%) and need for mechanical ventilation (16.9% vs 2.1%) (p-values <0.001). After propensity matching two well matched cohorts of 4140 patients were included in the analysis. Higher rates of mortality (15.3 v 4.8%), need for hospitalization (58.1% vs 25.1%) and mechanical ventilation (16.9% vs 3.9%) persisted in patients with TE, compared to patients without TE after propensity matching (p < 0.001).

Previous studies have reported 16% to 31% of COVID-19 patients to have some form of TE/complications and higher mortality in Covid-19 patients with TE.^{1,4} Our data from a very large sample of all-comers COVID-19 patients (including outpatients) shows a smaller prevalence of TE. This also provides more comprehensive estimates of the increased risk of worse outcomes including mortality in COVID-19 patients with TE events and underscores the need for intensive screening and thromboprophylaxis in these patients. Our study is limited because of not accounting for comorbidities / anticoagulation or anti-platelet therapy – however, it still represents a broad outline of the burden of TE in the largest sample of COVID-19 patients so far reported.

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- Bilaloglu S, Aphinyanaphongs Y, Jones S, Iturrate E, Hochman J, Berger JS. Thrombosis in hospitalized patients with COVID-19 in a New York City health system. *JAMA* 2020;324:799– 801. https://doi.org/10.1001/jama.2020.13372. PMID:32702090; PMCID: PMC7372509.
- Al-Samkari H, Karp Leaf RS, Dzik WH, Carlson JCT, Fogerty AE, Waheed A, Goodarzi K, Bendapudi PK, Bornikova L, Gupta S, Leaf DE, Kuter DJ, Rosovsky RP. COVID-19 and

coagulation: bleeding and thrombotic manifestations of SARS-CoV-2 infection. *Blood* 2020; 136:489–500. https://doi.org/10.1182/blood. 2020006520. PMID:32492712; PMCID: PMC7378457.

 Pakhchanian H, Raiker R, Mukherjee A, Khan A, Singh S, Chatterjee A. Outcomes of COVID-19 in CKD patients: a multicenter electronic medical record cohort study. *Clin J Am Soc Nephrol* 2021. https://doi.org/10.2215/ CJN.13820820. CJN.13820820 Online ahead of print.

 Mondal S, Quintili AL, Karamchandani K, Bose S. Thromboembolic disease in COVID-19 patients: a brief narrative review. *J Intensive Care* 2020;8:70. https://doi.org/10.1186/ s40560-020-00483-y. PMID:32939266; PMCID: PMC7487447.

https://doi.org/10.1016/j.amjcard.2021.03.003