COVID-19 infection in multiple myeloma patients - retrospective analysis of 371 Czech patients -

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IV. INTERNÍ HEMATOLOGICKÁ KLINIKA FAKULTNÍ NEMOCNICE HRADEC KRÁLOVÉ





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Disclosures

Company	Relationship
BMS	Medical Advisory Board Member, Travel expenses
Janssen	Medical Advisory Board Member
GSK	Medical Advisory Board Member
Amgen	Medical Advisory Board Member, Travel expenses
Sanofi	Medical Advisory Board Member, Travel expenses

(none related to this presentation)





Introduction



Myeloma population is vulnerable to infections and patients are at high risk of death of any infection



SARS-CoV-2, a "novel" RNA virus caused pandemic in erarly spring 2020 Causes COVID-19 disease with variable clinical course and outcome



Outcomes of immunocompromised patients infected with COVID-19 are poor The risk factors associated with death are not well understood



No established therapy for severe disease exists





EXACT Lancet Oncol. 2020;21:1309, Lancet 2020; 395:497., Clin Infect Dis. 2009;49(8):1211.

AI	Europe	North America	Asia	South Amer	ica Africa	a Oceania								
#	Country, Other 1	Total Cases ↓†	New Cases ↓†	Total Deaths ↓†	New Deaths ↓†	Total Recovered ↓↑	New Recovered ↓↑	Active Cases 1	Serious, Critical \downarrow†	Tot Cases/ 1M pop ↓↑	Deaths/ 1M pop ↓	Total Tests ↓†	Tests/ 1M pop ↓↑	Population 1
1	<u>Peru</u>	2,156,451		198,568		N/A	N/A	N/A	1,159	64,344	5,925	17,026,824	508,041	33,514,653
2	<u>Hungary</u>	814,064	+246	30,077	+3	778,401	+143	5,586	26	84,523	3,123	6,651,684	690,633	9,631,290
3	<u>Bosnia and</u> <u>Herzegovina</u>	219,010	+930	9,951	+32	192,218		16,841		67,250	3,056	1,162,083	356,836	3,256,632
4	<u>North</u> Macedonia	181,620	+701	6,153	+24	161,515	+1,119	13,952		87,180	2,954	1,203,819	577,850	2,083,273
5	<u>Gibraltar</u>	5,402	+3	97		5,233	+6	72		160,397	2,880	343,517	10,199,739	33,679
6	Montenegro	119,602	+533	1,780	+6	110,016	+696	7,806	11	190,401	2,834	649,041	1,033,242	628,160
7	Czechia	1,681,681	+588	30,408		1,648,424	+13	2,849	14	156,690	2,833	36,581,788	3,408,504	10,732,505
	14-day COVID-19 case notification rate													





IV. IHK

https://koronavirus.mzcr.cz/, https://www.ecdc.europa.eu/en/covid-19/situation-updates/weekly-maps-coordinated-restriction-free-movement

Patients and methods



- Retrospective analysis of patients with multiple myeloma and COVID-19 infection
- Patients who were diagnosed with COVID-19 between March 2020 and May 2021 were included in analysis
- Data were extracted and analyzed via Czech RMG registry (Registry of Monoclonal Gammopathies (<u>https://rmg.healthregistry.org/</u>), all patients provide informed consent prior to registry entry





Patients characteristics

	Total (N=371)		
Sex	n=371		
Man	198 (53.4%)		
Woman	173 (46.6%)		
ECOG at diagnosis MM	n=369		
PS 0	100 (27.1%)		
PS 1	173 (46.9%)		
PS 2	65 (17.6%)		
PS 3–4	31 (8.4%)		
ISS at diagnosis MM	n=359		
Stage 1	130 (36.2%)		
Stage 2	103 (28.7%)		
Stage 3	126 (35.1%)		
M-protein type at diagnosis MM	n=368		
IgG	233 (63.3%)		
IgA	65 (17.7%)		
LC only	61 (16.6%)		
Biclonal	4 (1.1%)		
IgD	2 (0.5%)		
IgM	3 (0.8%)		
Light chain type at diagnosis MM	n=368		
Карра	242 (65.8%)		
Lambda	122 (33.2%)		
Biclonal	4 (1.1%)		
Immunoparesis at diagnosis MM	n=284		
immunoparesis of 1 Ig	64 (22.5%)		
immunoparesis of 2 Ig	170 (59.9%)		

Treatment patterns	Total (N=371)
Number of tr. lines before COVID-19	
mean (sd)	1.3 (1.3)
median (min.–max.)	1 (0–7)
Without MM treatment before COVID-19	22 (6.3%)
Transplantation before COVID-19	136 (38.7%)
IMiD before COVID-19	195 (55.6%)
PI before COVID-19	238 (67.8%)
MoAB before COVID-19	30 (8.5%)

COVID-19 during MM treatment	197	53.1%
COVID-19 patients currently without MM treatment	13	3.5%
COVID-19 before 1st MM treatment line	9	2.4%
COVID-19 between MM treatment lines	9	2.4%
COVID-19 after currently last MM treatment line	123	33.2%





Course of pandemia in Czech MM population







Symptoms and course of infection

Symptoms

Proportion of patients (%)

Asymptomatic	18.6%	81.4%
Fever > 37 °C	49.6%	50.4%
Cough	39.1%	60.9%
Shortness of breath or difficulty breathing	35.0%	65.0%
Muscle or joint pains	10.0%	90.0%
Headache	8.4%	91.6%
New loss of taste or smell	7.8%	92.2%
Chills	3.8%	96.2%
Nausea/vomiting	3.8%	96.2%
Diarrhoea	3.0%	97.0%
Sore throat	1.3%	98.7%
"Covid toe" (itchy rash on toes)	0.5%	99.5%
Neurologic symptoms	0.5%	99.5%
Neuropathy	0.0%	100.0%
Other*	5.4%	94.6%
0	% 25% 50% 3	75% 100%

*Other: fatigue, anorexia, weakness, stroke, cramps, hallucination, cardiac failure, hematemesis, pain of skin, rhinitis, tremor

Severity and hospitalization (n=167, 45.0%)



Care type





Mild – mild symptoms without pneumonia , **Moderate** – respiratory symptoms + pneumonia at CT, **Severe** – respiratory rate \geq 30, oxygen saturation \leq 93%, oxygenation index \leq 300 **Critically severe** – respiratory failure with mechanical ventilation, shock, other organ failure

Proportion of patients (%)

Respiratory support and treatment

Respiratory support combination



Treatment





* Other: most frequently antibiotics and corticosteroids, Bamlanivimab, Isoprinosine, paralen, G-CSF...

Outcome



	Ν	mean (SD)	median (min–max)
Length of positivity (days)	79	23 (16)	20 (1–84)

Patient status to last update in registry







Association of selected variables with outcome

	Total (N=371)	Survivors (N=279)	Non-survivors (N=92)	p-value
Age at COVID-19 diagnosis (years)	n=353	n=266	n=87	0.0003
mean (sd)	67.1 (10.4)	65.8 (10.5)	70.8 (9.2)	
median (min.–max.)	69.0 (37.0–91.0)	68.0 (37.0–91.0)	72.0 (45.0–91.0)	
Immunoparesis at diagnosis MM	n=284	n=213	n=71	0.4060
immunoparesis of 1 Ig	64 (22.5%)	46 (21.6%)	18 (25.4%)	0.5151
immunoparesis of 2 Ig	170 (59.9%)	132 (62.0%)	38 (53.5%)	0.2122
Number of tr. lines before COVID-19	n=351	n=265	n=86	0.7090
mean (sd)	1.3 (1.3)	1.3 (1.2)	1.5 (1.6)	
median (min.–max.)	1 (0–7)	1 (0–7)	1 (0–7)	
Treatment patterns				
Without MM treatment before COVID-19	22 (6.3%)	19 (7.2%)	3 (3.5%)	0.3080
Transplantation before COVID-19	136 (38.7%)	113 (42.6%)	23 (26.7%)	0.0110
IMiD before COVID-19	195 (55.6%)	154 (58.1%)	41 (47.7%)	0.1050
PI before COVID-19	238 (67.8%)	184 (69.4%)	54 (62.8%)	0.2880
MoAB before COVID-19	30 (8.5%)	18 (6.8%)	12 (14.0%)	0.0470
Comorbidities				
Obesity (BMI > 30)	33 (10.4%)	25 (10.5%)	8 (10.3%)	1.0000





Association of selected variables with outcome

	Total (N=371)	Survivors (N=279)	Non-survivors (N=92)	p-value
Severity	n=301	n=218	n=83	< 0.001
Mild	175 (58.1%)	160 (73.4%)	15 (18.1%)	< 0.001
Moderate	60 (19.9%)	43 (19.7%)	17 (20.5%)	0.8729
Severe	34 (11.3%)	12 (5.5%)	22 (26.5%)	< 0.001
Critically severe	32 (10.6%)	3 (1.4%)	29 (34.9%)	< 0.001
Type of hospitalization	n=331	n=248	n=83	< 0.001
Without hospitalization	176 (53.2%)	169 (68.1%)	7 (8.4%)	< 0.001
Standard department	90 (27.2%)	61 (24.6%)	29 (34.9%)	0.0866
Intensive care unit	65 (19.6%)	18 (7.3%)	47 (56.6%)	< 0.001
Respiratory support	n=310	n=227	n=83	
AIRVO or non-invasive LV	48 (15.5%)	12 (5.3%)	36 (43.4%)	< 0.001
ALV	18 (5.8%)	4 (1.8%)	14 (16.9%)	< 0.001
Oxygen	92 (29.7%)	42 (18.5%)	50 (60.2%)	< 0.001





Conclusion



Myeloma patients are at high risk of death during or after COVID-19



¼ of infected patients die directly or as a consequence of COVID-19



1/2 of patients requiring oxygen die, 3/4 of patients requiring NIPV/ALV die



40% of patients pretreated with MoABs die





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