Critique and Debate

Was Wuhan the early epicenter of the COVID-19 pandemic? – A critique

Recently, Worobey et al. (2022) published a report with the title "The Huanan Seafood Wholesale Market in Wuhan was the early epicenter of the COVID-19 pandemic" that succinctly summarizes their study [1]. A pre-print version of this study had earlier elicited a series of high-profile media coverages [2, 3]. All these reports deliver a social-political message that the Huanan market is the epicenter of COVID-19.

At the outset, we shall clarify the meaning of "epicenter" in the context of the pandemic. Epicenter, narrowly defined, is the point on the earth's surface directly above an earthquake. Hence, in this context, epicenter should be the place from which the pathogen spread globally to cause the pandemic. By this definition, what Worobey et al. identified is not the epicenter of the global pandemic for the obvious reason that the conclusion is based entirely on SARS-CoV-2 samples collected in Wuhan. Where else could they have pinpointed the epicenter with their samples? Had they used only virus data from the Antarctica, they would have concluded that some place on that continent is the epicenter. A relevant example is the island province across the Taiwan Strait. Studies that analyzed the virus data entirely from the island have also shown that the Taoyuan International airport south of the city of Taipei is the "epicenter". In short, what Worobey et al. (2022) shows is that the early epidemic of Wuhan centered around its seafood market, analogous to the Taoyuan airport "epicenter" of Taiwan.

In a technical sense, Worobey et al.'s title is vague. Did they simply mean that the Huanan Market is the "epicenter" of the early *local* epidemic in Wuhan? Nevertheless, the juxtaposition of "epicenter" and "pandemic" in their title must have meant the global epicenter. Indeed, the global media has read Worobey et al. to mean Wuhan with its seafood market is the epicenter of the entire pandemic [2-6].

Worobey et al. have failed to follow the standard practice of presenting a scientific report in the context of previous publications on the same subject. This is particularly important when the conclusion is diametrically opposed to those of previous publications. Worobey et al. should have compared their conclusions with studies that have more extensive data geographically. Furthermore, although the interest is in the early phase of the pandemic, it has been shown that samples from the subsequent periods can be informative about the early phase by interpolation. After all, later samples are far more abundant and better organized when the human societies became aware of the onset of epidemics.

We have compiled a set of reports on viral samples from wild animals [7-15] which collectively are far more global by geography than the report we critique here. Another set of diverse studies provides evidence that SARS-CoV-2 may have been spreading worldwide weeks or even months prior to the epidemic in Wuhan in December 2019 [16-23]. Such reports have been brushed aside due to a mis-conception on the onset of epidemics.

The misconception is most explicitly stated in a recent news report [24] as follows: "the idea of a pandemic origin outside China is preposterous to many scientists, because there's simply no way SARS-CoV-2 could have come from some foreign place to Wuhan and triggered an explosive outbreak there without first racing through humans at the site of its origin." In the absence of an evolutionary perspective, that human and chimpanzee could have a common ancestor would be equally "preposterous". Cohen's other points including the integrity of Chinese scientists is not worthy of a response.

Using the branching process to model the evolution of epidemics, Ruan et al. [25, 26] and Kucharski et al. [27] have shown that invasions into a new population could trigger local epidemics only sporadically in the early phase. Local epidemics may even reach an alarming level of infections before fading out on its own. In this sense, there may be many earlier local epidemics (or endemics) that rise and fall before the eventual success to spread globally from the true epicenter of the pandemic. The many reports of local infections prior to the global pandemic could be such manifestations[16-23].

Ruan et al. [28]'s title "The twin-beginnings of COVID-19 in Asia and Europe—one prevails quickly" may be a most explicit analysis of the multiple early events. Their analysis has corroborated the earlier sampling results from the Lombardy region of Italy [19, 20]. All these studies have concluded that Wuhan is not likely to be the epicenter of the COVID-19 pandemic. Finally, however the critiques and debates in the scientific community may resolve the issue, it is unfortunate that the pre-print version of Worobey et al. has attained the unwarranted publicity on a subject of enormous social and political implications. The pre-print platform should be used by the scientific community to speed up exchanges, rather than by investigators to influence the societies at large before being debated among scientists.

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