A Framework for Federated Analysis of Health Data Using Multiparty Homomorphic Encryption

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Abstract: Although federated data analysis represents a significant contribution toward ensuring data privacy, the risk

of information leakage from the intermediate results exchanged during the analysis process still exists. These risks become even more emphasised when analysing sensitive data such as health records. One of the approaches to mitigate these issues is homomorphic encryption, a novel encryption algorithm which allows for performing computations over encrypted data. This article presents a federated data analysis framework where intermediate analysis results are exchanged and processed as ciphertexts and where data sources are connected in a decentralised manner by forming multiple clusters, with each cluster having a central node. Besides processing encrypted information, another advantage of the homomorphic encryption algorithms is the support for a multiparty encryption scheme. A workflow for creating a shared public and evaluation key is presented, where central nodes are part of the workflow and data sources only receive the shared keys. Furthermore, as data analysis examples, workflows for Kaplan-Meier survival analysis and distributed means a shared public and evaluation that the process of the state of

value are presented, whose results do match those obtained through centralized analysis. As a last step of the federated data analysis, multiparty decryption of the final result occurs.

1 INTRODUCTION

In health data analysis, where datasets consist of patients' health records, numerous security and privacy requirements need to be met before sharing data for analysis purposes. Since health data is considered sensitive and is subject to strict data protection legislation, a secure and reliable framework that prevents data leakage and protects patient information is a prerequisite. In a European context, All EU member states have been introduced to the General Data Protection Regulation (GDPR) act on data protection and privacy, which defines, among others, the use of health data for patient treatment and research purposes. Still, these countries have additional data protection acts for the use of health data put into force (Hansen et al., 2021), which can hinder data sharing and make the legal framework challenging to interpret, especially in the case of cross-border data sharing. Federated data analysis is one of the approaches that can address the beforementioned challenges. The federated approach allows data analysis algorithms to be applied to remotely

located datasets without sharing data directly with the data analyst. Instead, intermediate analysis results are exchanged and combined with results from other data sources. However, it has been proven that intermediate results can be exploited to retrieve information on background data (Zhu et al., 2019; Yang et al., 2023). One approach towards mitigating these issues would be to exchange all information in an encrypted format by encoding data into ciphertexts. This would prevent access to plaintext information without the corresponding decryption key, while still allowing data computation.

The above-mentioned approach can be addressed by using homomorphic encryption (HE) (Naehrig et al., 2011), which is an encryption algorithm capable of performing arithmetic operations over encrypted data. The results remain encrypted with decryption being the last step of the computing process. The data can be therefore computed or analysed safely without the possibility of unauthorized access to data or the intermediate results. This makes it especially suitable for the analysis of sensitive information, such as patient health records. Besides healthcare, areas of

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application include cloud computing, finance, the Internet of Things (IoT), and more. Arithmetic operations supported are addition and multiplication, from which other operations, i.e. subtraction and division can be derived. Secure computation and exchange of information within the federated analysis framework is enabled through multi-party computing algorithms, where parties have to collectively compute the critical security functions such as shared encryption key generation or the distributed result decryption. Some of the challenges with applying HE include the computational overhead in data processing and the complexity of the data analysis algorithms that need to be rewritten for the efficient use of HE.

HE applications and privacy-preserving federated data analysis are central points of this research. An overview of the related work is given in section 2, followed by a description of the proposed federated analysis framework and HE algorithms used. Finally, data analysis examples with implementation and workflow descriptions are presented and peerreviewed research studies, primarily conducted using the centralized analysis approach, were successfully reproduced. The presented federated data analysis framework is derived from the use cases from the EU project ORCHESTRA (ORCHESTRA Cohort, n.d.) whose plan is to create a Pan-European cohort for the conduct of prospective and retrospective studies to improve the prevention and treatment of COVID-19.

2 RELATED WORK

Most federated data analysis frameworks are designed to operate in a star topology where all data sources share intermediate results with the central server. There are, however, reports on decentralised layouts, which are developed or proposed to address various issues during analysis, not necessarily related to data privacy or security. (Wainakh et al., 2020) investigates potential benefits of hierarchical federated learning, such as no direct access from the central server to the data providers. (Hosseinalipour et al., 2020) describes the term fog computing, an approach for performing a computation in a heterogeneous network environment to prevent network overload, where devices don't have to be connected to the central server and can provide computational resources. Moreover, parameter aggregation can be done on a middle layer before sending results to the central node in the case of federated learning. (Hosseinalipour et al., 2022) presents another example of a distributed machine

learning model to connect a large number of devices with various properties, such as limited uplink transmission and communication protocol. With the advent of HE, it became possible to perform basic arithmetic operations (e.g. addition, subtraction, multiplication) over the encrypted data. (Froelicher et al., 2021) reports using secure multi-party computation for conducting biomedical studies. The developed system requires all parties to be interconnected and is capable of exchanging and processing only encrypted information. A single party cannot decrypt the study's results at any point. In addition, the model has a collective key switching feature implemented, which allows results to be decrypted only by a querier's secret key. (Sav et al., 2021) is another example of the secure multi-party computation suitable for neural network training, which is network topology agnostic in the context that parties can be arranged into any network topology. Furthermore, all parties are foreseen as data owners whose intermediate updates exchanged during the training process remain encrypted. Because of the HE, the last two examples add computational overhead to the process, and adding a new server to the model requires adjustments for all parties. (Blatt et al., 2020) reports the application of HE on genomewide association studies. R DataSHIELD (Marcon et al., 2021) is a tool for privacy-preserving distributed data analysis, well known in the area of clinical data analysis. Statistical operations are performed at the data source, where a person or a group performing the analysis receives only summary parameters or a final result. From the network topology point of view, R DataSHIELD is designed to work in a star topology.

3 FEDERATED ANALYSIS FRAMEWORK

The proposed federated data analysis framework and the communication flow are depicted in Figure 1. The data sources are depicted as hospitals that communicate only with their central nodes, sharing only encrypted information with intermediate results. Furthermore, hospitals are isolated from the researcher as well as other central nodes and underlying hospitals. Thus, the researcher does not contact the hospitals directly. The framework reflects the data infrastructure from the ORCHESTRA project, where three national hubs (central nodes) provide access to data delivered by the underlying data sources. The number of data sources assigned to each national hub varies which has also been

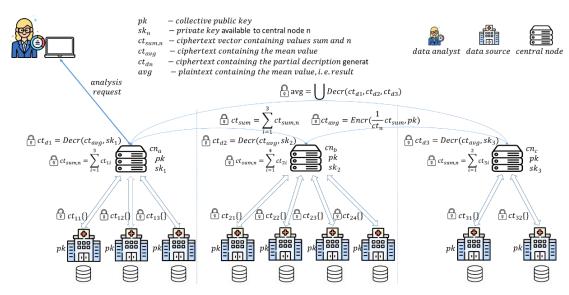


Figure 1: Federated data analysis framework with the communication workflow for computing the mean value.

considered in the presented framework. The advantage of applying HE algorithms is that the data sources only need to provide a connection to the central node (national hub) for handling encrypted data analysis requests without outsourcing their data and exposing the intermediate results that might cause sensitive information breaches.

The information between the clusters, as well as between the clusters and the data sources, is exchanged in an encrypted way using HE algorithms. Besides encrypted data processing, the latter allows multi-party computing where all involved parties collaborate in cryptographic processes, such as generating shared public keys and distributed decryption of a ciphertext. In this framework, central nodes collaborate to create the shared public keys using their respective secret key shares. Central nodes share public keys with their data sources for application to their data. The data never leaves the data source during the analysis. At the end of the analysis workflow, central nodes collaborate to decrypt the ciphertext containing the result.

Compared to the frameworks presented in (Froelicher et al., 2021) and (Sav et al., 2021), this framework differs in that data sources remain hidden behind their central nodes, with no direct interaction across clusters. Furthermore, adding or removing a data source to the framework affects only the involved central node and introduces no overhead associated with generating new shared homomorphic keys. Adding or removing a central node would affect the whole framework, which includes recreating the shared keys.

3.1 Homomorphic Encryption Workflow

The term crypto-context is used throughout the article, which denotes the selected encryption scheme and configured encryption parameters. The HE keys can be grouped into two categories: secret keys which are never shared with other nodes, and public keys which are shared with other nodes in the framework. The latter comprises a public key for encrypting the vector containing data, an evaluation rotation key for accessing the values at the specific index of the encrypted vector, and an evaluation multiplication key for reducing the amount of noise in the ciphertext caused by homomorphic multiplication operations.

The data analysis and the crypto-context generation are initiated from one place, e.g. by the researcher. A newly created crypto-context is sent to one of the central nodes, which acts as a first contact point and is also in charge of forwarding the crypto-context to the rest of the central nodes. This is followed by the message exchange between three central nodes with respective secret key shares sk_1 , sk_2 and sk_3 , who collaborate to create a public key pk, evaluation multiplication key mk and evaluation rotation key rk. These keys can be shared with data sources. Secret key shares on the other hand remain stored on corresponding central nodes.

The number of communication rounds for creating the shared public keys depends on the number of involved parties, i.e. central nodes. For the presented framework, up to 7 communication rounds per shared key are necessary. In particular, pk, rk and mk require n-1, n-1 and 2n-1 communication

rounds, respectively, where n is the number of involved parties. The data sources can now request shared keys from their central nodes. Distributed decryption requires all central nodes to collaborate and use their secret key shares sk_1 , sk_2 and sk_3 to compute the partial decryptions which are then joined into a final decryption, which is equivalent to the plaintext of an analysis result.

HE algorithms allow basic arithmetic operations, namely addition and multiplication. Subtraction can be derived from addition without applying a special algorithm. The division can be derived from multiplication by applying a special approximation algorithm. In the case of the presented framework, the Goldschmidt iterative division algorithm (Goldschmidt, 1964) as an approximation method is used, represented by the equation (1), where *x* is the number for which the reciprocal is computed:

$$\frac{1}{x} = \frac{1}{1 - (1 - x)} = \prod_{i=0}^{\infty} (1 + (1 - x)^{2^{i}})$$

$$\approx \prod_{i=0}^{d} (1 + (1 - x)^{2^{i}})$$
(1)

and where crypto-context parameters implicitly determine d through a maximum number of multiplication operations allowed.

The CKKS homomorphic encryption scheme for approximate arithmetic (Cheon et al., 2017) is chosen because it can be applied to real numbers and it supports multi-party computing. All HE operations used in the federated analysis workflows, including multi-party algorithms, are supported by the CKKS scheme and are implemented in the programming library used for the framework development (Section The crypto-context parameters (ciphertext dimension and modulus in particular) are selected to achieve the 128-bit level of security. The distributed decryption is part of the multi-party algorithms supported by the programming library. It is performed only by central nodes as a last step of the data analysis, where each node after computing partial decryption using their respective secret key shares, performs a fusion of the partial decryptions into a readable result ready to be shared with the data analyst.

Each data source ds_i provides a harmonised dataset in a CSV (Comma-Separated Values) format. Before starting an analysis, ds_i has to request shared keys pk and mk from the associated central node.

3.2 Data Analysis Workflows

3.2.1 Distributed Mean Value

Following is the workflow for finding the mean value of the analysed dataset:

- 1. Data source ds_i reads the CSV file with the individual patient information in a harmonised format, for which the mean value is computed.
- 2. Data source ds_i computes the local sum and encrypts the vector array of the result and the number of individuals with the pk. The created ciphertext ct_i is then sent to the corresponding central node.
- 3. Central nodes receive ct_i from the underlying data sources, after which a local sum $ct_{sum,i}$ and a local number of individuals $ct_{n,i}$ are computed by adding ct_i , followed by the addition between central nodes to find the overall sum ct_{sum} and the overall number of individuals ct_n . This step is done entirely using HE algorithms.
- 4. Since the average value is computed by multiplying the ct_{sum} with the reciprocal value of ct_n , the Goldschmidt division (1) is used, together with shared keys rk and mk. The ct_n is first scaled to a range of [0,2]. The maximum number of individuals must be set in advance to find the scaling factor. The Goldschmidt algorithm runs in seven iterations due to predefined parameters of the crypto context. Furthermore, seven iterations have proven to be sufficient for the accurate result. The result is then rescaled back.
- 5. The reciprocal value is multiplied by the sum to get the ciphertext ct_{avg} with the mean value.
- 6. Distributed decryption of ct_{avg} between central nodes takes place.

3.2.2 Survival Analysis

The survival analysis, in this case, is computed using the Kaplan-Meier estimator, which is presented by the equation (2):

$$\hat{S}(t) = \prod_{i,t_i \le T} \left(1 - \frac{d_i}{n_i}\right) \tag{2}$$

where t_i is a time when at least one event happened, d_i is the number of events that occurred at time t_i , and n_i is the number of individuals known to have survived up to time t_i .

Given that all data sources have a harmonised dataset in a CSV format where each patient is described with the following values: patient ID, time

and survival event, the workflow for the survival analysis is conducted as follows:

- 1. Data source ds_i reads the CSV file with the individual patient information used for the survival analysis.
- 2. Data source ds_i computes the new vector with time point, number of negative outcomes and number of censored events (for each time point). In addition, the vector also contains the number of patients present in the CSV file. The resulting vector is encrypted using pk. The created ciphertext ct_i is then sent to the central node.
- 3. Central nodes receive ct_i from the underlying data sources, after which they are added into one ciphertext ct_{res} .
- 4. Distributed decryption of *ct*_{res} between central nodes takes place.
- 5. This step is performed by the data analyst: values from the decrypted vector are used to compute the number of patients per time point and finally, the survival rate using the Kaplan-Meier estimator presented by (2). At this point, the result can be visualised.

The workflow for the survival analysis is similar to those described in (Froelicher et al., 2021).

4 EXPERIMENTAL SETUP

The HE part of the federated analysis framework is built using OpenFHE (Al Badawi et al., 2022), a C++ library that provides implementations of fully homomorphic encryption schemes. The rest of the framework is developed in C++ and with the ZeroMQ library (Hintjens, 2013) for message exchange between the nodes. The structure of the network messages that are sent between the parties comprises a header section for sorting the messages, and a body section containing the data to be sent. Furthermore, the experiments were conducted using the 'tc' tool in Linux to simulate a network latency of 20 ms and a bandwidth of 1 Gbit/s. The framework prototype was compiled and tested on a single machine with a Linux operating system and the following hardware: 2x AMD EPYC 7261 8 core processor, 125 GiB of working memory, and 128 GB of SSD storage. All parties in the analysis framework communicate using TCP/IP sockets. It is important to point out that OpenFHE already provides various approximation algorithms based on Chebyshev polynomials, such as a reciprocal or a sigmoid function. This paper however reports the implementation of the previously mentioned Goldschmidt algorithm for finding the reciprocal value of a given number. The data used for the analysis are coming from the study reported in (Samstein et al., 2019).

Central node and data source have server and client components. Before starting the shared keys generation or data analysis, server components should be up and running and listening to incoming connections. Client components initiate the outgoing connection based on the input received by the server component. The researcher is provided with an associated client component to establish a connection with one of the central nodes that serves as a main contact point, which in this case is a central node cn_a . Before execution of the analysis or HE keys generation, the availability of all the central nodes is checked.

5 RESULTS

Compared with the centralised and non-encryption analysis methods, the following processes need to be taken into concern: crypto-context distribution, HE keys generation, data encryption, distributed decryption, message exchange and homomorphic arithmetic operations. These require additional computation and storage resources, thus representing a time overhead. For example, better performance can be achieved by manually reducing noise in the ciphertext generated by multiplication operations. Both data analysis examples show that analysis data does not have to leave the data source, and all intermediate results are shared only as ciphertexts. Furthermore, communication with data sources is done only through central nodes, which makes them hidden from other parties in the framework. By using data from the study presented in (Samstein et al., 2019), the obtained results are identical to those obtained through a centralized analysis approach. Key generation operation includes serialising keys and saving them into separate files for later use. Subsequent data analysis operations therefore do not require key generation. The data analysis workflows were tested on 9 data sources unevenly distributed among 3 central nodes, as displayed in Figure 1. Each data source had access to a different number of data items, where the total number was 1662.

The workflow for computing the distributed mean value of the TMB (Tumor Mutational Burden) variable was tested first. Figure 2 shows the share of key generation, data analysis and distributed decryption per central node in the overall wall time. The analysis time overhead is mainly caused by the Goldschmidt algorithm and its loop composed of multiple homomorphic multiplication operations.

The algorithm is iterated seven times inside the loop. The obtained result is 11.9692 and it does match the one obtained through the centralized analysis approach.

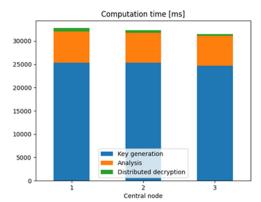


Figure 2: Wall time for the federated encrypted mean value analysis.

The same setup was used for testing the survival analysis workflow, the total number of data items or patient records was again 1662. The analysis was conducted on the same subset of data as in the original study. Figure 3 shows the comparison of the results between centralized and federated encrypted analysis, which do match. The share of key generation, data analysis and distributed decryption per central node in the overall wall time is presented in Figure 4. The measured wall time covers steps 1 to 4 from the survival analysis workflow. The wall time for step 5 was not measured, as it was not done by the data analyst after the distributed decryption.

The wall time for the distributed key generation is longer for the distributed mean value analysis due to the higher multiplicative depth setting in the cryptocontext configuration necessary to perform multiple instances of the homomorphic multiplication operation in the division algorithm. Survival analysis, on the other hand, does not require homomorphic multiplication operations.

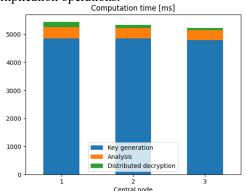


Figure 3: Wall time for the federated encrypted survival analysis.

6 CONCLUSION

This work presents a framework suitable for the federated analysis of sensitive health data. Decentralised network topology allows for creating trust domains composed of a central node and multiple data sources. Presented workflows for computing mean value and survival analysis using the Kaplan-Meier estimator demonstrate intermediate results do not have to be shared with other parties in plaintext format. Instead, they are encrypted and further computed using HE algorithms. The decryption process is done at the very end of the analysis process in a distributed way, requiring all central nodes to collaborate. Finally, the obtained centralized and federated encrypted analysis results are identical.

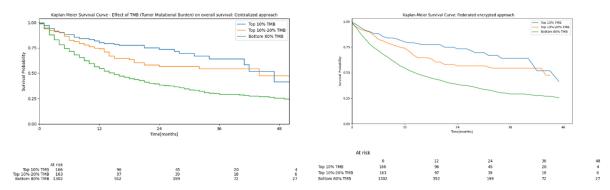


Figure 4: Survival analysis results comparison between centralized (left) and federated encrypted (right) approach.

7 FUTURE WORK

A preliminary step towards future work will be further optimisation of the HE crypto-context parameters that would reduce the analysis time overhead. Afterwards, alternatives to distributed decryption will be explored, such as key switching, where the result ciphertext gets re-encrypted so that it only can be decrypted with a secret key owned by the researcher. The final objective in the presented future work plan is the implementation of regression analysis workflow to support machine learning algorithms that are running in multiple epochs and iterations. This increases noise in the ciphertext due to HE multiplication, requiring ciphertext refreshing or bootstrapping (Micciancio & Polyakov, 2021) to reduce the amount of noise in the ciphertext and allow subsequent operations and decryption. In a multi-party setting, this activity has to be done collaboratively by all central nodes, which presents an additional step in the analysis workflow.

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